

=> e honda mitsuo/au

E1	70	HONDA MITSUNOBU/AU
E2	25	HONDA MITSUNORI/AU
E3	325 -->	HONDA MITSUO/AU
E4	329	HONDA MITSURU/AU
E5	10	HONDA MITSUTAKA/AU
E6	3	HONDA MITSUTATSU/AU
E7	19	HONDA MITSUTERU/AU
E8	59	HONDA MITSUTOSHI/AU
E9	3	HONDA MITSUYA/AU
E10	8	HONDA MITSUYASU/AU
E11	6	HONDA MITSUYO/AU
E12	9	HONDA MITSUYOSHI/AU

=> s e3 and bcg

L1 46 "HONDA MITSUO"/AU AND BCG

=> dup rem l1

PROCESSING COMPLETED FOR L1

L2 30 DUP REM L1 (16 DUPLICATES REMOVED)

=> d bib ab 1-

YOU HAVE REQUESTED DATA FROM 30 ANSWERS - CONTINUE? Y/(N):y

L2 ANSWER 1 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2006:512935 CAPLUS

DN 145:6612

TI A method of prime-boost vaccination for AIDS caused by HIV-1 CRF01\_AE strain

IN Honda, Mitsuo; Matsuo, Kazuhiro; Hamano, Takaichi; Izumi, Yasuyuki; Promkhatkaew, Duanthanorm; Balachandra, Kruavon; Sutthent, Ruengpung

PA Japan Science and Technology Agency, Japan; Japan as Represented by Director General of National Institute of Infectious Diseases; Departement of Medical Sciences, Ministry of Public Health, Thailand

SO PCT Int. Appl., 39 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2006057454	A1	20060601	WO 2005-JP22221	20051125
	W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, KE, KG, KM, KN, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
	RW:	AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM			
	JP 2006149234	A	20060615	JP 2004-341283	20041125

PRAI JP 2004-341283 A 20041125

AB An invention of this application is a method of prime-boost vaccination comprising a priming step by a recombinant BCG vaccine and one or more boosting steps by a recombinant vaccine, wherein both of the recombinant BCG vaccine for priming step and the recombinant vaccine for boosting steps have at least one gene of HIV-1 CRF01\_AE strain. In said invention, it is a preferred embodiment that both of the

recombinant vaccines have at least gag gene of HIV-1 CRFO1-AE strain. The present inventors have found that in the case of using recombinant BCG as a priming antigen in combination with other viral vector-based vaccine as a boosting antigen, quite efficiently enhanced cellular immune response could be induced against HIV-1 CRFO 1\_AE whereupon the present invention has been achieved.

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 2 OF 30 USPATFULL on STN

AN 2006:247197 USPATFULL

TI Recombination bcg vaccine

IN Honda, Mitsuo, Tokyo, JAPAN

Matsuo, Kazuhiro, Kanagawa, JAPAN

Kanekiyo, Masaru, Tokyo, JAPAN

PI US 2006210586 A1 20060921

AI US 2003-524586 A1 20030813 (10)

WO 2003-JP10303 20030813

20050331 PCT 371 date

PRAI JP 2002-237610 20020816

DT Utility

FS APPLICATION

LREP WENDEROTH, LIND & PONACK, L.L.P., 2033 K STREET N. W., SUITE 800,

WASHINGTON, DC, 20006-1021, US

CLMN Number of Claims: 2

ECL Exemplary Claim: 1

DRWN 5 Drawing Page(s)

LN.CNT 640

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A recombinant BCG vaccine being transformed with an expression vector that has a polynucleotide encoding a foreign antigenic protein, wherein the polynucleotide is a modified one in which a third position of each codon is substituted with G or C without a change of an amino acid. This recombinant BCG vaccine has an excellent expression rate of antigenic protein and, as a result, capable of inducing a sufficient immune response against target infectious disease, cancer, or the like at the same dose as that of the typical BCG vaccine.

L2 ANSWER 3 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2006:1086167 CAPLUS

DN 145:453348

TI Priming-boosting vaccination with recombinant Mycobacterium bovis bacillus Calmette-Guerin and a nonreplicating vaccinia virus recombinant leads to long-lasting and effective immunity. [Erratum to document cited in CA143:365230]

AU Ami, Yasushi; Izumi, Yasuyuki; Matsuo, Kazuhiro; Someya, Kenji; Kanekiyo, Masaru; Horibata, Shigeo; Yoshino, Naoto; Sakai, Koji; Shinohara, Katsuaki; Matsumoto, Sohichi; Yamada, Takeshi; Yamazaki, Shudo; Yamamoto, Naoki; Honda, Mitsuo

CS Division of Experimental Animal Research, AIDS Research Center, Division of Biosafety Control and Research, National Institute of Infectious Diseases, Toyama 1-23-1, Shinjuku-ku, Tokyo, 162-8640, Japan

SO Journal of Virology (2006), 80(20), 10288

CODEN: JOVIAM; ISSN: 0022-538X

PB American Society for Microbiology

DT Journal

LA English

AB On page 12873, Table 1, in groups 1, 2 and 3, rBCG was primed at week 3, followed by booster immunization of rDIs at weeks 50 and 54. In the same table, groups 4 and 5, rDIs was primed at weeks 0 and 12, followed by booster immunization of rBCG at week 50. Then all animals were challenged with virulent SHIV KS661c at week 57. On page 12874, Figure 2: The weeks after immunization shown on the x axis in panel A should read: "3, 7, 11, 19, 27, 35, 50, 53, 54, and 56.". The weeks after immunization shown on

the x axis in panel B should read: "0, 4, 8, 16, 24, 32, 50, 52, 53, 54, and 56.". Although the patterns and magnitudes of the kinetics were almost the same as the original ones, the standard deviation of the ELISPOT data at the peak response at 54 wk after immunization was 500, which was five times more than originally reported. Spot-forming cells were counted by using a KS ELISPOT system after 35 wk of immunization. Before that, we counted SFCs using an inverted microscope. On page 12878, Acknowledgments, paragraph 1, the last sentence of the paragraph should be deleted. These changes do not alter the conclusions of the article.

L2 ANSWER 4 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN  
AN 2006:1171430 CAPLUS  
DN 146:160902  
TI Strategy for development of prophylactic and therapeutic vaccines against HIV  
AU Matsuo, Kazuhiro; Yamamoto, Naoki; Honda, Mitsuo  
CS AIDS Research Center, National Institute of Infectious Diseases, Tokyo, Japan  
SO Igaku no Ayumi (2006), 218(10), 923-930  
CODEN: IGAYAY; ISSN: 0039-2359  
PB Ishiyaku Shuppan  
DT Journal; General Review  
LA Japanese  
AB A review discusses development of prophylactic and therapeutic vaccines such as DNA vaccine, BCG vaccine application and antigen peptides vaccine for HIV infection.

L2 ANSWER 5 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN  
DUPLICATE 1  
AN 2006:315079 BIOSIS  
DN PREV200600309141  
TI Intradermal and oral immunization with recombinant Mycobacterium bovis BCG expressing the simian immunodeficiency virus Gag protein induces long-lasting, antigen-specific immune responses in guinea pigs.  
AU Kawahara, Mamoru [Reprint Author]; Matsuo, Kazuhiro; Honda, Mitsuo  
CS Univ Occupat and Environm Hlth, Dept Biochem and Mol Pathophysiol, Sch Med, Yahatanishi Ku, 1-1 Iseigaoka, Kitakyushu, Fukuoka 8078555, Japan  
mamokawa@med.uoeh-u.ac.jp  
SO Clinical Immunology (Orlando), (APR 2006) Vol. 119, No. 1, pp. 67-78.  
ISSN: 1521-6616.  
DT Article  
LA English  
ED Entered STN: 14 Jun 2006  
Last Updated on STN: 14 Jun 2006  
AB To develop a new recombinant BCG (rBCG) vaccine, we constructed rBCG that expresses the full-length Gag protein of simian immunodeficiency virus (rBCG-SIVGag) at a level of 0.5 ng/mg after 3 weeks of bacterial cell culture. Intradermal (i.d.) inoculation of guinea pigs with 0.1 mg of rBCG-SIVGag resulted in the induction of delayed-type hypersensitivity (DTH) responses to both purified protein derivative (PPD) of tuberculin and SIV Gag p27 protein; responses that were maintained for the duration of the 50-week study. In contrast, guinea pigs orally vaccinated with 160 mg of the same antigen exhibited a long-lasting DTH response to the SIV Gag p27 protein, but mounted no response to PPD. Proliferative responses to SIV Gag p27 and PPD antigens were detected in both i.d. and orally immunized animals; however, the levels of PPD-specific responses were significantly higher in guinea pigs immunized by the i.d. than the oral route. A significant increase in the level of PPD- and SIV Gag p27-specific IFN gamma mRNA expression was also detected in both immunization groups receiving rBCG-SIVGag. In addition, both i.d. and oral immunization with rBCG-SIVGag induced PPD- and SIV Gag p27-specific serum IgG responses. Insertion of the SIV gag gene into BCG did not appear to change the ability of rBCG-immunized animals to elicit PPD-specific immune responses. These results indicate that rBCG-SIVGag

has the ability to effectively induce long-lasting, cell-mediated and humoral immunity against both viral and bacterial. antigens in guinea pigs, suggesting that rBCG-Gag has the potential to elicit immunities specific not only for tuberculosis but also for HIV at human doses. (c)  
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L2 ANSWER 6 OF 30 USPATFULL on STN  
AN 2005:143826 USPATFULL  
TI Radio lan access authentication system  
IN Honda, Mitsuo, Tokyo, JAPAN  
Matsuo, Kazuhiro, Kanagawa, JAPAN  
Izumi, Yasuyuki, Tokyo, JAPAN  
PI US 2005123561 A1 20050609  
AI US 2003-515253 A1 20021120 (10)  
WO 2002-JP12125 20021120  
PRAI JP 2002-145132 20020520  
DT Utility  
FS APPLICATION  
LREP WENDEROTH, LIND & PONACK, L.L.P., 2033 K STREET N. W., SUITE 800,  
WASHINGTON, DC, 20006-1021, US  
CLMN Number of Claims: 7  
ECL Exemplary Claim: 1  
DRWN 14 Drawing Page(s)  
LN.CNT 356

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB This invention provides a recombinant BCG vaccine transformed by an expression vector having polynucleotide coding for exogenous antigenic protein, characterized in that, the BCG vaccine is used for initial antigen stimulus in immune induction by plural antigen stimulations and also provides a method for the immune induction where the initial antigen stimulus is carried out by the said BCG vaccine and one or more additional antigenic stimulation(s) is/are carried out by non-BCG vaccine expressing the same antigenic protein.

L2 ANSWER 7 OF 30 USPATFULL on STN  
AN 2005:62585 USPATFULL  
TI Recombinant vaccinia virus vaccine  
IN Honda, Mitsuo, Tokyo, JAPAN  
Matsuo, Kazuhiro, Kanagawa, JAPAN  
Ohsu, Takeaki, Kanagawa, JAPAN  
Miyamura, Tatsuo, Tokyo, JAPAN  
Matsuura, Yoshiharu, Osaka, JAPAN  
Ishii, Koji, Tokyo, JAPAN  
Kato, Kenzo, Chiba, JAPAN  
PI US 2005053620 A1 20050310  
AI US 2004-495974 A1 20040903 (10)  
WO 2001-JP10141 20011120  
DT Utility  
FS APPLICATION  
LREP WENDEROTH, LIND & PONACK, L.L.P., 2033 K STREET N. W., SUITE 800,  
WASHINGTON, DC, 20006-1021  
CLMN Number of Claims: 19  
ECL Exemplary Claim: 1  
DRWN 7 Drawing Page(s)  
LN.CNT 849

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention provides a recombinant vaccinia virus strain DIs that possesses a polynucleotide encoding a foreign antigenic protein in the non-essential gene region of the chromosome DNA and expresses the antigenic protein; and provides a highly-safe, vaccinia virus vaccine containing the recombinant virus strain DIs as the active ingredient. The invention also provides a method of using the vaccinia virus strain DIs as a vector for protein expression.

L2 ANSWER 8 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN  
 DUPLICATE 2  
 AN 2006:8301 BIOSIS  
 DN PREV200600008996  
 TI Priming-boosting vaccination with recombinant Mycobacterium bovis bacillus  
 Calmette-Guerin and a nonreplicating vaccinia virus recombinant leads to  
 long-lasting and effective immunity.  
 AU Ami, Yasushi; Izumi, Yasuyuki; Matsuo, Kazuhiro; Someya, Kenji; Kanekiyo,  
 Masaru; Horibata, Shigeo; Yoshino, Naoto; Sakai, Koji; Shinohara,  
 Katsuaki; Matsumoto, Sohichi; Yamada, Takeshi; Yamazaki, Shudo; Yamamoto,  
 Naoki; Honda, Mitsuo [Reprint Author]  
 CS Natl Inst Infect Dis, Ctr AIDS Res, Shinjuku Ku, Toyama 1-23-1, Tokyo  
 1628640, Japan  
 mhonda@nih.go.jp  
 SO Journal of Virology, (OCT 2005) Vol. 79, No. 20, pp. 12871-12879.  
 CODEN: JOVIAM. ISSN: 0022-538X.  
 DT Article  
 LA English  
 ED Entered STN: 14 Dec 2005  
 Last Updated on STN: 14 Dec 2005  
 AB .Virus-specific T-cell responses can limit immunodeficiency virus type 1  
 (HIV-1) transmission and prevent disease progression and so could serve as  
 the basis for an affordable, safe, and effective vaccine in humans. To  
 assess their potential for a vaccine, we used Mycobacterium bovis bacillus  
 Calmette-Guerin (BCG)-Tokyo and a replication-deficient vaccinia  
 virus strain (DIs) as vectors to express full-length gag from simian  
 immunodeficiency viruses (SIVs) (rBCG-SIVgag and rDisSIVgag). Cynomolgus  
 macaques were vaccinated with either rBCG-SIVgag dermally as a single  
 modality or in combination with rDisSIVgag intravenously. When  
 cynomolgus macaques were primed with rBCG-SIVgag and then boosted with  
 rDisSIVgag, high levels of gamma interferon (IFN-gamma) spot-forming cells  
 specific for SIV Gag were induced. This combination regimen elicited  
 effective protective immunity against mucosal challenge with pathogenic  
 simian-human immunodeficiency virus for the 1 year the macaques were under  
 observation. Antigen-specific intracellular IFN-gamma activity was  
 similarly induced in each of the macaques with the priming-boosting  
 regimen. Other groups receiving the opposite combination or the  
 single-modality vaccines were not effectively protected. These results  
 suggest that a recombinant M. bovis BCG-based vector may have  
 potential as an HIV/AIDS vaccine when administered in combination with a  
 replication-deficient vaccinia virus DIs vector in a priming-boosting  
 strategy.

L2 ANSWER 9 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN  
 DUPLICATE 3  
 AN 2005:399376 BIOSIS  
 DN PREV200510190449  
 TI Mycobacterial codon optimization enhances antigen expression and  
 virus-specific immune responses in recombinant Mycobacterium bovis bacille  
 Calmette-Guerin expressing human immunodeficiency virus type 1 Gag.  
 AU Kanekiyo, Masaru; Matsuo, Kazuhiro; Hamatake, Makiko; Hamano, Takaichi;  
 Ohsu, Takeaki; Matsumoto, Sohichi; Yamada, Takeshi; Yamazaki, Shudo;  
 Hasegawa, Atsuhiko; Yamamoto, Naoki; Honda, Mitsuo [Reprint  
 Author]  
 CS Natl Inst Infect Dis, AIDS Res Ctr, Shinjuku Ku, 1-23-1 Toyama, Tokyo  
 1628640, Japan  
 mhonda@nih.go.jp  
 SO Journal of Virology, (JUL 2005) Vol. 79, No. 14, pp. 8716-8723.  
 CODEN: JOVIAM. ISSN: 0022-538X.  
 DT Article  
 LA English  
 ED Entered STN: 5 Oct 2005  
 Last Updated on STN: 5 Oct 2005

AB Although its potential for vaccine development is already known, the introduction of recombinant human immunodeficiency virus (HIV) genes to *Mycobacterium bovis* bacille Calmette-Guerin (BCG) has thus far elicited only limited responses. In order to improve the expression levels, we optimized the codon usage of the HIV type 1 (HIV-1) p24 antigen gene of gag (p24 gag) and established a codon-optimized recombinant BCG (rBCG)-p24 Gag which expressed a 40-fold-higher level of p24 Gag than did that of nonoptimized rBCG-p24 Gag. Inoculation of mice with the codon-optimized rBCG-p24 Gag elicited effective immunity, as evidenced by virus-specific lymphocyte proliferation, gamma interferon ELISPOT cell induction, and antibody production. In contrast, inoculation of animals with the nonoptimized rBCG-p24 Gag induced only low levels of immune responses. Furthermore, a dose as small as 0.01 mg of the codon-optimized rBCG per animal proved capable of eliciting immune responses, suggesting that even low doses of a codon-optimized rBCG-based vaccine could effectively elicit HIV-1-specific immune responses.

L2 ANSWER 10 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 4

AN 2005:114055 CAPLUS

DN 142:334776

TI Vaccination of rhesus macaques with recombinant *Mycobacterium bovis* bacillus Calmette-Guerin Env V3 elicits neutralizing antibody-mediated protection against simian-human immunodeficiency virus with a homologous but not a heterologous V3 motif

AU Someya, Kenji; Cecilia, Dayaraj; Ami, Yasushi; Nakasone, Tadashi; Matsuo, Kazuhiro; Burda, Sherri; Yamamoto, Hiroshi; Yoshino, Naoto; Kaizu, Masahiko; Ando, Shuji; Okuda, Kenji; Zolla-Pazner, Susan; Yamazaki, Shudo; Yamamoto, Naoki; Honda, Mitsuo

CS AIDS Research Center, National Institute of Infectious Diseases, Tokyo, 162-8640, Japan

SO Journal of Virology (2005), 79(3), 1452-1462

CODEN: JOVIAM; ISSN: 0022-538X

PB American Society for Microbiology

DT Journal

LA English

AB Although the correlates of vaccine-induced protection against human immunodeficiency virus type 1 (HIV-1) are not fully known, it is presumed that neutralizing antibodies (NAb) play a role in controlling virus infection. In this study, we examined immune responses elicited in rhesus macaques following vaccination with recombinant *Mycobacterium bovis* bacillus Calmette-Guerin expressing an HIV-1 Env V3 antigen (rBCG Env V3). We also determined the effect of vaccination on protection against challenge with either a simian-human immunodeficiency virus (SHIV-MN) or a highly pathogenic SHIV strain (SHIV-89.6PD). Immunization with rBCG Env V3 elicited significant levels of NAb for the 24 wk tested that were predominantly HIV-1 type specific. Sera from the immunized macaques neutralized primary HIV-1 isolates in vitro, including HIV-1BZ167/X4, HIV-1SF2/X4, HIV-1C12/X4, and, to a lesser extent, HIV-1MNp/X4, all of which contain a V3 sequence homologous to that of rBCG Env V3. In contrast, neutralization was not observed against HIV-1SF33/X4, which has a heterologous V3 sequence, nor was it found against primary HIV-1 R5 isolates from either clade A or B. Furthermore, the viral load in the vaccinated macaques was significantly reduced following low-dose challenge with SHIV-MN, and early plasma viremia was markedly decreased after high-dose SHIV-MN challenge. In contrast, replication of pathogenic SHIV-89.6PD was not affected by vaccination in any of the macaques. Thus, we have shown that immunization with an rBCG Env V3 vaccine elicits a strong, type-specific V3 NAb response in rhesus macaques. While this response was not sufficient to provide protection against a pathogenic SHIV challenge, it was able to significantly reduce the viral load in macaques following challenge with a nonpathogenic SHIV. These observations suggest that rBCG vectors have the potential to deliver an appropriate virus immunogen for desirable immune elicitation.

RE.CNT 54 THERE ARE 54 CITED REFERENCES AVAILABLE FOR THIS RECORD

## ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 11 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
STN . DUPLICATE 5  
AN 2005:338939 BIOSIS  
DN PREV200510120020  
TI Specific immune response and pathological findings in BALB/c mice  
inoculated with recombinant BCG expressing HIV-1 antigen.  
AU Wiriyarat, Witthawat; Sukpanichnant, Sanya; Sittisombut, Nopporn;  
Balachandra, Kruavon; Promkhatkaew, Duanthanorm; Butraporn, Raywadee;  
Sutthent, Ruengpung; Boonlong, Jotika; Matsuo, Kazuhiro; Honda,  
Mitsuo; Warachit, Paijit; Puthavathana, Pilaipan [Reprint Author]  
CS Mahidol Univ, Siriraj Hosp, Fac Med, Dept Microbiol, Bangkok 10700,  
Thailand  
siput@mahidol.ac.th  
SO Asian Pacific Journal of Allergy and Immunology, (MAR 2005) Vol. 23, No.  
1, pp. 41-51.  
ISSN: 0125-877X.  
DT Article  
LA English  
ED Entered STN: 31 Aug 2005  
Last Updated on STN: 31 Aug 2005  
AB Recombinant BCGs (rBCGs) containing extrachromosomal plasmids with  
different HIV-1 insert sequences: nef, env (V3J1 and E9Q), gag p17 or  
whole gag p55 were evaluated for their immunogenicity, safety and  
persistent infection in BALB/c mice. Animal injected with, rBCG-pIJKV3J1,  
rBCG-pSO gag p17 or rBCG-pSO gag p55 could elicit lymphocyte proliferation  
as tested by specific HIV-1 peptides or protein antigen. Inoculation with  
various concentration of rBCG-pSO gag p55 generated satisfactory specific  
lymphocyte proliferation in dose escalation trials. The rBCG-pSO gag p55  
recovered from spleen tissues at different time interval post-inoculation  
could express the HIV protein as determined by ELISA p24 antigen detection  
kit. This result indicated that the extrachromosomal plasmid was stable  
and capable to express Gag protein. It was also demonstrated that rBCGs  
did not cause serious pathological change in the inoculated animals. The  
present study suggested the role of BCG as a potential vehicle  
for using in HIV vaccine development.

L2 ANSWER 12 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN  
AN 2004:162604 CAPLUS  
DN 140:216160  
TI Recombinant BCG vaccines against infection, cancer and other  
diseases  
IN Honda; Mitsuo; Matsuo, Kazuhiro; Kanekiyo, Masaru  
PA Japan Science and Technology Corporation, Japan; Japan as Represented by  
Director General of National Institute of Infectious Diseases  
SO PCT Int. Appl., 31 pp.  
CODEN: PIXXD2

DT Patent  
LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2004016280	A1	20040226	WO 2003-JP10303	20030813
	W: IN, JP, US				
	RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR				
	EP 1535627	A1	20050601	EP 2003-788098	20030813
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, SK				
	US 2006210586	A1	20060921	US 2005-524586	20050331
PRAI	JP 2002-237610	A	20020816		
	WO 2003-JP10303	W	20030813		
AB	A recombinant BCG vaccine obtained by transformation with an				

expression vector carrying a polynucleotide coding for an endemic antigenic protein, which recombinant BCG vaccine consists of a modified type polynucleotide comprising a polynucleotide having the third base of each of the codons thereof substituted with G or C without changing of the type of amino acid. This recombinant BCG vaccine excels in the amount of antigenic protein expressed, so that even with the same dosage as employed for conventional BCG vaccines, the recombinant BCG vaccine can induce satisfactory immune response to target infectious diseases, cancer, etc. In example, recombinant BCG comprising hsp60 gene and HIV-1 gag p24 gene was prepared for use as vaccine.

RE.CNT 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 13 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
STN DUPLICATE 6  
AN 2004:450507 BIOSIS  
DN PREV200400452713  
TI The normalization of guinea pig leukocyte fractions and lymphocyte subsets  
in blood and lymphoid tissues using a flow cytometric procedure.  
AU Takizawa, Mari; Chiba, Jo; Haga, Shinji; Asano, Toshihiko; Yamamoto,  
Naoki; Honda, Mitsuo [Reprint Author]  
CS AIDS Res CtrShinjuku Ku, Natl Inst Infect Dis, 1-23-1 Toyama, Tokyo,  
1628640, Japan  
SO Experimental Animals (Tokyo), (July 2004) Vol. 53, No. 4, pp. 321-329.  
print.  
ISSN: 1341-1357 (ISSN print).  
DT Article  
LA English  
ED Entered STN: 24 Nov 2004  
Last Updated on STN: 24 Nov 2004  
AB Many hematological and immunological parameters remain unclear in the  
study of the guinea pig. In this study, we established the mean values of  
blood counts, the percentage of leukocyte fractions and lymphocyte subsets  
in blood and various lymphoid tissues of the guinea pig with a flow  
cytometric procedure using MIL4/SSC. The mean counts of WBC and RBC in  
the blood were lower, and MCV and MCH were higher than those of other  
rodents, resembling those of humans. Furthermore, the mean percentages of  
blood lymphocytes were smaller and that of granulocyte was larger than  
those of other rodents, resembling those of humans. We further  
established a flow cytometric procedure for lymphocyte subsets and  
clarified the mean percentages of T- and B-cells, CD4+-, CD8+ and MHC  
Class II+- T-cells, and CD4-CD8- T-cells. The latter were morphologically  
larger in cell size and cytoplasm than CD4+- plus CD8+ T-cells, and this  
subset had a significantly higher percentage in newborn animals.  
Furthermore, the appearance of the MHC Class II+ T-cell subset was  
suggested to be a marker of hyper-activation of T-cells in BCG  
-immunized animals. Thus, both the novel flow cytometric procedure for  
leukocyte fractions and lymphocyte subsets, and the established normal  
values will be useful tools in studying guinea pigs as models of various  
diseases and biological phenomena.

L2 ANSWER 14 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN  
AN 2003:931202 CAPLUS  
DN 139:394884  
TI BCG vector and non-BCG vector encoding antigenic  
protein for use as vaccine and booster vaccine against infection and  
cancer  
IN Honda, Mitsuo; Matsuo, Kazuhiro; Izumi, Yasuyuki  
PA Japan Science and Technology Corporation, Japan; Japan as Represented by  
Director General of National Institute of Infectious Diseases  
SO PCT Int. Appl., 35 pp.  
CODEN: PIXXD2  
DT Patent



LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
	-----	----	-----	-----	-----
PI	WO 2003097087	A1	20031127	WO 2002-JP12125	20021120
	W: CA, JP, US				
	CA 2494359	A1	20031127	CA 2002-2494359	20021120
	US 2005123561	A1	20050609	US 2003-515253	20021120
	IN 2003DE00709	A	20050311	IN 2003-DE709	20030520
PRAI	JP 2002-145132	A	20020520		
	WO 2002-JP12125	W	20021120		

AB A recombinant BCG vaccine transformed by an expression vector which carries a polynucleotide encoding a foreign antigenic protein characterized by being usable for priming in immune induction via twice or more antigenic stimulations; and an immune induction method characterized by comprising priming with the use of the above BCG vaccine and boosting once or more with the use of a non-BCG vaccine expressing the same antigenic protein. Recombinant BCG (rBCG-SIVgag) and recombinant vaccinia Dis (rDis-SIVgaga) encoding SIV gag gene were prepared and tested.

RE.CNT 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 15 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
STN DUPLICATE 7

AN 2003:37096 BIOSIS

DN PREV200300037096

TI Dynamics of gamma interferon, interleukin-12 (IL-12), IL-10, and transforming growth factor beta mRNA expression in primary Mycobacterium bovis BCG infection in guinea pigs measured by a real-time fluorogenic reverse transcription-PCR assay.

AU Kawahara, Mamoru; Nakasone, Tadashi; Honda, Mitsuo [Reprint Author]

CS National Institute of Infectious Diseases, 1-23-1 Toyama, Shinjuku-ku, Tokyo, 162-8640, Japan  
mhonda@nih.go.jp

SO Infection and Immunity, (December 2002) Vol. 70, No. 12, pp. 6614-6620.  
print.

ISSN: 0019-9567 (ISSN print).

DT Article

LA English

ED Entered STN: 8 Jan 2003

Last Updated on STN: 8 Jan 2003

AB The guinea pig has been utilized as a model for studying infectious diseases because its reactions closely resemble those of humans biologically and immunologically. However, the cytokine responses in this animal remain to be studied. Initially, we established a quantitative assay using a real-time reverse transcription-PCR (RT-PCR) to measure guinea pig gamma interferon (IFN-gamma), interleukin-12 (IL-12), IL-10, and transforming growth factor beta (TGF-beta) mRNA. By preparing primer-fluorogenic probe sets for these cytokines and standard RNA templates corresponding to the target sequence of each cytokine, we obtained linear standard curves essential for quantitative determination. In guinea pigs immunized by intradermal (i.d.) vaccination with the Tokyo strain of Mycobacterium bovis BCG (0.1 mg) or else hyperimmunized with the same vaccine (10 mg) given intravenously (i.v.), peripheral blood mononuclear cells (PBMCs) at 4 weeks showed an increase in IFN-gamma mRNA expression in the latter but not the former animals. However, at week 10, IFN-gamma mRNA expression was markedly elevated in PBMCs, spleen cells, and cells in bronchoalveolar lavage fluid in both the i.d.- and the i.v.-immunized animals, the level of expression being 10 times higher in the latter. In contrast, the expression levels of IL-12 mRNA in PBMCs, spleen cells, and BAL cells were not enhanced in either group at 10 weeks postimmunization. The expression of IL-10 and TGF-beta

increased slightly only in PBMCs. Regardless of differences in the levels of cytokine responses, the magnitudes of the purified protein derivative of tuberculin-specific delayed-type hypersensitivity (DTH) skin reactions for the two groups did not differ significantly at 8 weeks postvaccination. In this study, we quantitatively measured IL-10, IL-12, TGF-beta, and IFN-gamma mRNA in BCG-immunized guinea pigs and showed that the level of IFN-gamma mRNA expression does not necessarily reflect the magnitude of the DTH response, suggesting that there may be an intricate relationship between protective immunity, the level of IFN-gamma, and the DTH response. Thus, our quantitative assay would be of use for the development of vaccines using guinea pig models.

L2 ANSWER 16 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
STN DUPLICATE 8

AN 2003:63648 BIOSIS

DN PREV200300063648

TI Oral recombinant Mycobacterium bovis bacillus Calmette-Guerin expressing HIV-1 antigens as a freeze-dried vaccine induces long-term, HIV-specific mucosal and systemic immunity.

AU Kawahara, Mamoru; Hashimoto, Akira; Toida, Ichiro; Honda, Mitsuo [Reprint Author]

CS AIDS Research Center, National Institute of Infectious Diseases, 1-23-1 Toyama, Shinjuku-ku, Tokyo, 162-8640, Japan  
mhonda@nih.go.jp

SO Clinical Immunology (Orlando), (December 2002) Vol. 105, No. 3, pp. 326-331. print.  
ISSN: 1521-6616 (ISSN print).

DT Article

LA English

ED Entered STN: 22 Jan 2003  
Last Updated on STN: 22 Jan 2003

AB Induction of HIV-1-specific immune responses was evaluated using a recombinant BCG (rBCG) vector-based vaccine expressing HIV-1 Env V3 peptide (rBCG-pSOV3J1). rBCG-pSOV3J1 was manufactured as a freeze-dried preparation based on good laboratory practice guidelines. Guinea pigs were immunized with the freeze-dried rBCG vaccine by oral administration to test the effectiveness of what is generally considered the most convenient and practical route for vaccination. While delayed-type hypersensitivity (DTH) skin reactions to purified protein derivative were not detected in any of the animals receiving oral rBCG-pSOV3J1, HIV-1 V3J1 antigen-specific DTH responses were detected in all of the immunized guinea pigs 1.5 years after immunization. In addition, significant proliferative responses against HIV-1 V3J1 antigen were measured in peripheral blood mononuclear cells and splenocytes from all animals receiving oral rBCG. Interestingly, intestinal intraepithelial lymphocytes from the animals also exhibited high levels of proliferative activity against HIV-1 V3J1 antigen. These results suggest that oral vaccination of guinea pigs with freeze-dried rBCG-pSOV3J1 induces high levels of functional T cells specific for HIV-1 antigens in both mucosal and systemic compartments and suggest that this approach has potential for use as a vaccine against HIV-1.

L2 ANSWER 17 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
STN DUPLICATE 9

AN 2003:19497 BIOSIS

DN PREV200300019497

TI Combined intrarectal/intradermal inoculation of recombinant Mycobacterium bovis bacillus Calmette-Guerin (BCG) induces enhanced immune responses against the inserted HIV-1 V3 antigen.

AU Kawahara, Mamoru; Matsuo, Kazuhiro; Nakasone, Tadashi; Hiroi, Takachika; Kiyono, Hiroshi; Matsumoto, Sohkiichi; Yamada, Takeshi; Yamamoto, Naoki; Honda, Mitsuo [Reprint Author]

CS AIDS Research Center, National Institute of Infectious Diseases, 1-23-1 Toyama, Shinjuku-ku, Tokyo, 162-8640, Japan

mhonda@nih.go.jp

SO Vaccine, (13 December 2002) Vol. 21, No. 3-4, pp. 158-166. print.  
ISSN: 0264-410X (ISSN print).

DT Article

LA English

ED Entered STN: 1 Jan 2003  
Last Updated on STN: 1 Jan 2003

AB The development of a successful recombinant Mycobacterium bovis bacillus Calmette-Guerin (rBCG) vector-based vaccine for human immunodeficiency virus type 1 (HIV-1) requires the induction of high levels of HIV-1-specific immunity while at the same time maintaining immunity to tuberculosis. To examine a combined vaccination strategy for enhancement of immune responses specific for HIV-1, guinea pigs were inoculated with either a single or combination intradermal (i.d.), intrarectal (i.r.) and intranasal (i.n.) administration of rBCG-pSOV3J1 which secretes a chimeric protein of HIV-1 V3J1 peptide and alpha-antigen. Significant level of delayed-type hypersensitivity to both V3J1 peptide and tuberculin was induced in guinea pigs inoculated with human doses of rBCG-pSOV3J1 by a combination of intrarectal and intradermal routes. Guinea pigs inoculated by combined routes also had significantly higher titers of HIV-1-specific serum IgG and IgA compared with those animals immunized only intrarectally, which led to the enhanced neutralization activity against HIV-1MN. In addition, the induction of high levels of IFNgamma and interleukin-2 (IL-2) mRNA in PBMC, splenocytes, and intraepithelial lymphocytes from the immunized animals was detected until at least 110 weeks post-inoculation. These results suggest that enhanced immune responses specific for HIV-1 are efficiently induced by combined intrarectal and intradermal immunization with rBCG-HIV, and antigen-specific Th1-type memory cells are maintained for more than 2 years in the immunized animals. Thus, inoculation with rBCG-HIV by combined routes represents an effective vaccination strategy to elicit high levels of HIV-1-specific immune responses.

L2 ANSWER 18 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
STN DUPLICATE 10

AN 2002:438698 BIOSIS

DN PREV200200438698

TI Selective expansion of perforin-positive CD8+ T cells by immature dendritic cells infected with live Bacillus Calmette-Guerin mycobacteria.

AU Tsunetsugu-Yokota, Yasuko [Reprint author]; Tamura, Hideto; Tachibana, Mikiko; Ogata, Kiyoyuki; Honda, Mitsuo; Takemori, Toshitada

CS Department of Immunology, National Institute of Infectious Diseases, 1-23-1, Toyama-cho, Shinjuku-ku, Tokyo, 162-8640, Japan  
yyokota@nih.go.jp

SO Journal of Leukocyte Biology, (July, 2002) Vol. 72, No. 1, pp. 115-124. print.  
CODEN: JLBIE7. ISSN: 0741-5400.

DT Article

LA English

ED Entered STN: 14 Aug 2002  
Last Updated on STN: 14 Aug 2002

AB Live, but not dead Bacillus Calmette-Guerin (BCG) is partially protective against infection by Mycobacterium tuberculosis, which causes a disease with high mortality in immune compromised individuals. We have shown that uptake of BCG induces maturation of immature dendritic cells (DCs) regardless of the viability of the bacteria. Importantly, when T cells are cocultured with live BCG-infected DCs, the proportion of CD45RA- perforin+ CD8+ T cells is markedly expanded markedly; however, little expansion is seen when T cells are cocultured with DCs harboring heat-killed BCG. The direct contact of T cells with live BCG-infected DCs was required for the expansion of perforin+ CD8+ T cells. These CD8+ T cells demonstrated a high level of killing activity against BCG-infected macrophages. There was little contribution of cytokines, including IFN-gamma, TNF-alpha, and

IL-12, to the expansion of CD8+ T cells by live BCG-infected DCs. We found that the interaction between BCG-infected DCs and CD8+ T cells through CD40/CD40L was crucial for the expansion and maturation of CD8+ T cells, the process of which was CD4-independent. In contrast, blocking the CD58/CD2 but not the CD40/CD40L interaction reduced production of IFN-gamma without affecting the maturation of CD8+ T cells. This indicates that the production of IFN-gamma and perforin by CD8+ T cells is mediated by distinct signals delivered from BCG-infected DCs. Thus, BCG-specific CD8+ CTL memory cells may be maintained for a long period of time in BCG-vaccinated hosts, and these cells could mature rapidly into effectors through the potent antigen-presenting function of DCs upon mycobacterial infection.

L2 ANSWER 19 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN  
 AN 2001:832585 CAPLUS  
 DN 136:117006  
 TI HIV mucosal vaccine: nasal immunization with rBCG-V3J1 induces a long term V3J1 peptide-specific neutralizing immunity in Th1- and Th2-deficient conditions  
 AU Hiroi, Takachika; Goto, Hironobu; Someya, Kenji; Yanagita, Manabu; Honda, Mitsuo; Yamanaka, Noboru; Kiyono, Hiroshi  
 CS Department of Mucosal Immunology, Research Institute for Microbial Diseases, Osaka University, Osaka, Japan  
 SO Journal of Immunology (2001), 167(10), 5862-5867  
 CODEN: JOIMA3; ISSN: 0022-1767  
 PB American Association of Immunologists  
 DT Journal  
 LA English  
 AB In the vaccine strategy against HIV, bacillus Calmette-Guérin (BCG), a live attenuated strain of Mycobacterium bovis, is considered to be one of potential vectors for mucosal delivery of vaccine Ag. We analyzed the induction of the Ag-specific Ab response by nasal immunization with recombinant BCG vector-based vaccine (rBCG-V3J1) that can secrete the V3 principal neutralizing epitope of HIV. Mice were nasally immunized with rBCG-V3J1 (10 µg) three times at weekly intervals. Four weeks after the initial immunization, high titers of V3J1-specific IgG Abs were seen in serum. These high levels of HIV-specific serum IgG responses were maintained for > 12 mo following nasal immunization without any booster immunization. V3J1-specific IgG-producing cells were detected in mononuclear cells isolated from spleen, nasal cavity, and salivary gland of the nasally vaccinated mice. Nasal rBCG-V3J1 also induced high levels of prolonged HIV-specific serum IgG responses in Th1 (IFN-γ-/-) or Th2 (IL-4-/-)-immunodeficient mice. Further, IgG3 was highest among V3 peptide-specific IgG subclass Ab responses in these immunodeficient mice as well as in wild-type mice. In addition, this Ag-specific serum IgG Abs induced by nasal immunization with rBCG-V3J1 possessed the ability to neutralize clin. isolate of HIV in vitro. These results suggested that the nasal rBCG-V3J1 system might be used as a therapeutic vaccine in addition to a prophylaxis vaccine for the control of AIDS.

RE.CNT 35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L2 ANSWER 20 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN  
 AN 2002:89940 BIOSIS  
 DN PREV200200089940  
 TI Cross-clade neutralizing antibody production against human immunodeficiency virus type 1 clade E and B' strains by recombinant Mycobacterium bovis BCG-based candidate vaccine.  
 AU Chujoh, Yoshitomo; Matsuo, Kazuhiro; Yoshizaki, Hitomi; Nakasatomi, Tetsuya; Someya, Kenji; Okamoto, Yukari; Naganawa, Satoshi; Haga, Shinji; Yoshikura, Hiroshi; Yamazaki, Akihiro; Yamazaki, Shudo; Honda, Mitsuo [Reprint author]  
 CS AIDS Research Center, National Institute of Infectious Diseases, Toyama

1-23-1, Shinjuku-ku, Tokyo, 162-8640, Japan

mhonda@nih.go.jp

SO Vaccine, (12 December, 2001) Vol. 20, No. 5-6, pp. 797-804. print.  
CODEN: VACCDE. ISSN: 0264-410X.

DT Article

LA English

ED Entered STN: 24 Jan 2002

Last Updated on STN: 25 Feb 2002

AB The recombinant Mycobacterium bovis BCG (rBCG) vector-based vaccine secreting the V3 principal neutralizing epitope of human immunodeficiency virus type 1 (HIV-1) Japanese strain was reported to induce both humoral and cellular immune responses effectively (Proc. Natl. Acad. Sci. USA. 92 (1995) 10693). The antigen-secreting rBCG system was applied to the V3 epitope of clade E HIV-1 in this study. The V3 sequence of 19 amino acids (aa) and 15aa fused with mycobacterial alpha-antigen was not secreted while 12aa and 11aa sequences were successfully secreted from BCG cells. Serum IgG from guinea pig which was immunized with 12aa epitope-secreting recombinant BCG neutralized the WHO reference strain as well as primary field isolates of clade E virus. The serum IgG could also neutralize Thai B (clade B') strains which possessed a conserved GPGQ motif in their V3 sequences. These data suggest that the rBCG construct secreting the 12aa epitope is implicated in the development of a prophylactic vaccine in Thailand in which both clade E and B' viruses are prevalent.

L2 ANSWER 21 OF 30 USPATFULL on STN

AN 1999:36712 USPATFULL

TI Anti-AIDS secretory recombinant BCG vaccine

IN Matsuo, Kazuhiro, Kawasaki, Japan

Chujo, Yoshitomo, Kawasaki, Japan

Yamazaki, Akihiro, Kawasaki, Japan

Honda, Mitsuo, Mitaka, Japan

Yamazaki, Shudo, Higashiyamato, Japan

Tasaka, Hiromichi, Kure, Japan

PA Ajinomoto Co., Inc., Tokyo, Japan (non-U.S. corporation)

Japan as represented by Director General of Agency of National Institute of Health, Tokyo, Japan (non-U.S. government)

PI US 5885580 19990323

AI US 1997-972089 19971117 (8)

RLI Division of Ser. No. US 1996-619512, filed on 29 Mar 1996, now abandoned

PRAI JP 1994-178462 19940729

DT Utility

FS Granted

EXNAM Primary Examiner: Stucker, Jeffrey; Assistant Examiner: Park, Hankyel T.

LREP Oblon, Spivak, McClelland, Maier & Neustadt, P.C.

CLMN Number of Claims: 26

ECL Exemplary Claim: 1

DRWN 15 Drawing Figure(s); 11 Drawing Page(s)

LN.CNT 1556

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A vaccine containing Mycobacterium bovis BCG which secretes a fusion protein to be obtained by inserting a foreign antigen peptide into the molecular surface of a secretory protein, a carrier, having a signal peptide. BCG constituting the present invention secretes a fusion protein to be obtained by inserting a foreign antigen peptide into the molecular surface of an  $\alpha$ -antigen derived from mycobacteria. Said fusion protein has significantly increased antigenicity and immunogenicity. Therefore, when it is inoculated into animals, it is efficiently recognized by B cells which recognize said antigen, thereby effectively inducing the production of an antibody to said antigen. When said BCG itself is inoculated into animals, it continuously secretes said fusion protein in the bodies of the animals while continuously propagating therein. Therefore, said BCG is an extremely useful vaccine.

L2 ANSWER 22 OF 30 USPATFULL on STN  
 AN 1999:4042 USPATFULL  
 TI Anti-acids secretory recombinant BCG vaccine  
 IN Matsuo, Kazuhiro, Kawasaki, Japan  
 Chujo, Yoshitomo, Kawasaki, Japan  
 Yamazaki, Akihiro, Kawasaki, Japan  
 Honda, Mitsuo, Mitaka, Japan  
 Yamazaki, Shudo, Higashiyamato, Japan  
 Tasaka, Hiromichi, Kure, Japan  
 PA Ajinomoto Co., Inc., Tokyo, Japan (non-U.S. corporation)  
 Japan as represented by Director General of Agency of National Institute  
 of Health, Tokyo, Japan (non-U.S. corporation)  
 PI US 5858369 19990112  
 AI US 1997-975699 19971121 (8)  
 RLI Continuation of Ser. No. US 1996-619512, filed on 29 Mar 1996, now  
 abandoned  
 PRAI JP 1994-178462 19940729  
 DT Utility  
 FS Granted  
 EXNAM Primary Examiner: Stucker, Jeffrey; Assistant Examiner: Park, Hankyel T.  
 LREP Oblon, Spivak, McClelland, Maier & Neustadt, P.C.  
 CLMN Number of Claims: 16  
 ECL Exemplary Claim: 1  
 DRWN 15 Drawing Figure(s); 11 Drawing Page(s)  
 LN.CNT 1514

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A vaccine containing Mycobacterium bovis BCG which secretes a  
 fusion protein to be obtained by inserting a foreign antigen peptide  
 into the molecular surface of a secretory protein, a carrier, having a  
 signal peptide. BCG constituting the present invention  
 secretes a fusion protein to be obtained by inserting a foreign antigen  
 peptide into the molecular surface of an  $\alpha$ -antigen derived from  
 mycobacteria. Said fusion protein has significantly increased  
 antigenicity and immunogenicity. Therefore, when it is inoculated into  
 animals, it is efficiently recognized by B cells which recognize said  
 antigen, thereby effectively inducing the production of an antibody to  
 said antigen. When said BCG itself is inoculated into animals,  
 it continuously secretes said fusion protein in the bodies of the  
 animals while continuously propagating therein. Therefore, said  
 BCG is an extremely useful vaccine.

L2 ANSWER 23 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN  
 AN 1997:310334 CAPLUS  
 DN 126:304686  
 TI Analysis of the principal neutralization site of HIV-1 and vaccine  
 research  
 AU Okamoto, Yukari; Honda, Mitsuo  
 CS AIDS Res. Cent., Natl. Inst. Health, Tokyo, 162, Japan  
 SO Tanpakushitsu Kakusan Koso (1997), 42(7), 1201-1208  
 CODEN: TAKKAJ; ISSN: 0039-9450  
 PB Kyoritsu  
 DT Journal; General Review  
 LA Japanese  
 AB A review, with 20 refs., on variety of the neutralization sites of HIV-1  
 and their structure, modeling for 3-dimensional structure of HIV-1 PND  
 (principal neutralization determinant/domain), structure of HIV-1 vaccine  
 rBCG-V3J1 prepared by using the Japanese consensus sequence of HIV-1  
 inserted into the gene of BCG secretory  $\alpha$  antigen, and  
 neutralizing activity of serum IgG from guinea pigs immunized with the  
 vaccine. Anal. of prospective HIV-1 antigens useful for vaccine  
 development and drug design of anti-HIV-1 agents are also discussed.

L2 ANSWER 24 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1997:130690 CAPLUS  
 DN 126:223933  
 TI Vaccine therapy for AIDS  
 AU Someya, Kenji; Honda, Mitsuo  
 CS AIDS Research Center, National Institute of Health, Japan  
 SO Rinsho Kagaku (Osaka) (1997), 33(1), 49-56  
 CODEN: RIKAE; ISSN: 0385-0323  
 PB Esuato K. K.  
 DT Journal; General Review  
 LA Japanese  
 AB A review with 10 refs., on difficulties in developing vaccine for AIDS, vaccine that induces artificial immunity and its enhancement, anti-HIV vaccine antigen gp160, gp120, and gp41, second generation vaccine, preparation and evaluation of recombinant BCG vaccine, administration, and 3rd phase clin. test.

L2 ANSWER 25 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN  
 AN 1996:340573 CAPLUS  
 DN 125:8464  
 TI Anti-AIDS secretory recombinant BCG vaccine containing  $\alpha$ -antigen of acid-fast bacterium to enhance antigenicity  
 IN Matsuo, Kazuhiro; Chujo, Yoshitomo; Yamazaki, Akihiro; Honda, Mitsuo; Yamazaki, Shudo; Tasaka, Hiromichi  
 PA Ajinomoto Co., Inc., Japan; Japan, Agency of National Institute of Health  
 SO PCT Int. Appl., 56 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA Japanese  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9604009	A1	19960215	WO 1995-JP1515	19950731
	W: CN, JP, KR, US				
	RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
	CN 1136280	A	19961120	CN 1995-190959	19950731
	EP 745386	A1	19961204	EP 1995-926523	19950731
	EP 745386	B1	20040204		
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE				
	AT 258986	T	20040215	AT 1995-926523	19950731
	US 5885580	A	19990323	US 1997-972089	19971117
	US 5858369	A	19990112	US 1997-975699	19971121
PRAI	JP 1994-178462	A	19940729		
	WO 1995-JP1515	W	19950731		
	US 1996-619512	B3	19960329		

AB A vaccine containing Mycobacterium bovis BCG which secretes a fused protein obtained by inserting a foreign antigen peptide to the surface of a mol. of a secreted protein having a signal peptide as a carrier. The BCG secretes a fused protein obtained by inserting a foreign antigen peptide to the surface of a mol. of an  $\alpha$ -antigen originating in an acid-fast bacterium. The fused protein has extremely enhanced antigenicity and can efficiently induce the production of an antibody against the  $\alpha$ -antigen in animals. When the BCG can serve as a live vaccine; it continues to grow and secrete the fused protein in the animal body. Preparation of recombinant secretion vectors encoding fusion protein containing  $\alpha$ -antigen and the epitope of V3 surface antigen of HIV-1 was shown and their antigenicity were demonstrated in vivo. The antibodies thus produced also cross-react to V3 epitopes of types Thai-A and Thai-B.

L2 ANSWER 26 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN  
 AN 1996:574191 CAPLUS  
 DN 125:272948  
 TI Protective immune responses induced by recombinant Mycobacterium bovis bacillus Calmette-Guerin vector-based vaccine for HIV-1

AU Okamoto, Yukari; Ando, Shuji; Honda, Mitsuo  
CS AIDS Res. Cent., Natl. Inst. Health, Tokyo, 162, Japan  
SO Molecular Medicine (Tokyo) (1996), 33(Suppl. 447), 206-213  
CODEN: MOLMEL; ISSN: 0918-6557  
PB Nakayama Shoten  
DT Journal; General Review  
LA Japanese  
AB A review, with 13 refs., on the construction of human immunodeficiency virus (HIV) vaccine gene using HIV V3 PND region (V3J1) and recombinant BCG (Mycobacterium bacillus Calmette-Guerin), induction of HIV-PND specific delayed-type hypersensitivity (DTH) in guinea pig, and neutralizing activity of IgG produced by the vaccination. Difference of the neutralizing activity between activated peripheral blood monocyctic cells (PBMC) and resting PBMC is pointed out as a problems for evaluation of the neutralizing activity. The maintenance of Th1 function in acquired immunodeficiency syndrome (AIDS) is important, and the vaccine would contribute to it. The results of the vaccine are discussed using severe combined immunodeficiency (SCID) mouse and cynomolgus monkey.

L2 ANSWER 27 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN  
DUPLICATE 12  
AN 1996:21760 BIOSIS  
DN PREV199698593895  
TI Protective immune responses induced by secretion of a chimeric soluble protein from a recombinant Mycobacterium bovis bacillus Calmette-Guerin vector candidate vaccine for human immunodeficiency virus type 1 in small animals.

AU Honda, Mitsuo [Reprint author]; Matsuo, Kazuhiro; Nakasone, Tadashi; Okamoto, Yukari; Yoshizaki, Hitomi; Kitamura, Katsuhiko; Sugiura, Wataru; Watanabe, Kuhomi; Fukushima, Yoshiko; Haga, Shinji; Katsura, Yoshimoto; Tasaka, Hiromichi; Komuro, Katsutoshi; Yamada, Takeshi; Asan, Toshihiko; Yamazaki, Akihiro; Yamazaki, Shudo  
SO Proceedings of the National Academy of Sciences of the United States of America, (1995) Vol. 92, No. 23, pp. 10693-10697.  
CODEN: PNASA6. ISSN: 0027-8424.  
DT Article  
LA English  
ED Entered STN: 12 Jan 1996  
Last Updated on STN: 12 Jan 1996

AB A recombinant Mycobacterium bovis bacillus Calmette-Guerin (BCG) vector-based vaccine that secretes the V3 principal neutralizing epitope of human immunodeficiency virus (HIV) could induce immune response to the epitope and prevent the viral infection. By using the Japanese consensus sequence of HIV-1, we successfully constructed chimeric protein secretion vectors by selecting an appropriate insertion site of a carrier protein and established the principal neutralizing determinant (PND)-peptide secretion system in BCG: The recombinant BCG (rBCG)-inoculated guinea pigs were initially screened by delayed-type hypersensitivity (DTH) skin reactions to the PND peptide, followed by passive transfer of the DTH by the systemic route. Further, immunization of mice with the rBCG resulted in induction of cytotoxic T lymphocytes. The guinea pig immune antisera showed elevated titers to the PND peptide and neutralized HIV-MN, and administration of serum IgG from the vaccinated guinea pigs was effective in completely blocking the HIV infection in thymus/liver transplanted severe combined immunodeficiency (SCID)/hu or SCID/PBL mice. In addition, the immune serum IgG was shown to neutralize primary field isolates of HIV that match the neutralizing sequence motif by a peripheral blood mononuclear cell-based virus neutralization assay. The data support the idea that the antigen-secreting rBCG system can be used as a tool for development of HIV vaccines.

L2 ANSWER 28 OF 30 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN



AN 1994:466100 BIOSIS  
 DN PREV199497479100  
 TI HIV-PND-inserted secretory protein system of recombinant BCG  
 vector can effectively induce immune response to Japanese consensus HIV.  
 AU Honda, Mitsuo [Reprint author]; Matsuo, K.; Kitamura, K.;  
 Nakasone, T.; Okamoto, Y.; Watanabe, K.; Yoshizaki, H.; Fukushima, Y.;  
 Sugiura, W.; Tasaka, H.; Yamazaki, A.; Moritsugu, Y.; Yamada, K.;  
 Yamazaki, S.  
 CS AIDS Res. Cent., Tokyo, Japan.  
 SO TENTH INTERNATIONAL CONFERENCE ON AIDS, INTERNATIONAL CONFERENCE ON STD.  
 (1994) pp. 1) 74. Tenth International Conference on AIDS and the  
 International Conference on STD, Vol. 1; The global challenge of AIDS:  
 Together for the future.  
 Publisher: Tenth International Conference on AIDS, Yokohama, Japan.  
 Meeting Info.: Meeting. Yokohama, Japan. August 7-12, 1994.  
 DT Conference; (Meeting)  
 Conference; Abstract; (Meeting Abstract)  
 LA English  
 ED Entered STN: 31 Oct 1994  
 Last Updated on STN: 31 Oct 1994

L2 ANSWER 29 OF 30 CAPLUS COPYRIGHT 2007 ACS on STN  
 AN 1993:5257 CAPLUS  
 DN 118:5257  
 TI Purification of MPB70 and production of specific monoclonal antibodies  
 AU Haga, Shinji; Nakagawa, Masaro; Nagai, Sadamu; Miura, Kaoru; Honda,  
 Mitsuo  
 CS Dep. Cell. Immunol., Natl. Inst. Health, Tokyo, 141, Japan  
 SO Hybridoma (1992), 11(4), 483-92  
 CODEN: HYBRDY; ISSN: 0272-457X  
 DT Journal  
 LA English  
 AB MPB70 is a protein secreted into the culture filtrate of Mycobacterium  
 bovis BCG (substrain Tokyo 172), which is able to induce a  
 delayed-type hypersensitivity (DTH) skin reaction in guinea pigs immunized  
 with BCG-Tokyo. By high-pressure chromatofocusing and  
 size-exclusion HPLC, a further purified MPB70 protein was obtained, which  
 was visualized as a single band with a mol. mass of 22 kDa by SDS-PAGE. A  
 series of hybridoma cell lines that produced monoclonal antibodies (MAbs)  
 against the purified MPB70 protein was prepared, and 3 MAbs, Bov-1-3, with  
 strong antigen-binding capacities were established. Bov-1 was the most  
 potent MAb among them and binds to only a 22 kDa protein band in culture  
 filtrates of M. bovis, but not to bands in those of M. tuberculosis by  
 Western immunoblotting anal., suggesting that Bov-1 recognize a different  
 epitope of MPB70 from MAbs that have been shown previously to recognize  
 several species of mols. in culture filtrates of M. bovis. The purified  
 MPB70 protein elicited a strong DTH skin reaction in guinea pigs  
 sensitized with BCG-Tokyo vaccine. Bov-1 had no inhibitory  
 effect on generation of the DTH skin reaction, showing that MAb bound to  
 an epitope distinct from that inducing the skin reaction. All of the MAbs  
 were specific to MPB70 and each recognized a different epitope on MPB70.  
 MPB70 was not detected in the culture filtrate of M. tuberculosis H37Rv.

L2 ANSWER 30 OF 30 JAPIO (C) 2007 JPO on STN  
 AN 2006-149234 JAPIO  
 TI PRIME-BOOST VACCINATION METHOD  
 IN HONDA MITSUO; MATSUO KAZUHIRO; HAMANO RYUICHI; IZUMI YASUYUKI;  
 PROMKHATKAEW DUANTHANORM; BALACHANDRA KRUAUVON; SUTTHENT RUENG PUNG  
 PA JAPAN SCIENCE & TECHNOLOGY AGENCY  
 THAILAND MINISTRY OF PUBLIC HEALTH DEPARTMENT OF MEDICAL SCIENCES  
 NATIONAL INSTITUTE OF INFECTIOUS DISEASES  
 PI JP 2006149234 A 20060615 Heisei  
 AI JP 2004-341283 (JP2004341283 Heisei) 20041125  
 PRAI JP 2004-341283 20041125

SO PATENT ABSTRACTS OF JAPAN (CD-ROM), Unexamined Applications, Vol. 2006  
AB PROBLEM TO BE SOLVED: To provide a new vaccination strategy to HIV-1  
CEFO1AE.

SOLUTION: The prime-boost vaccination method comprises a priming step with a recombinant BCG vaccine and one or more boosting steps with the recombinant vaccine. Both of the recombinant BCG vaccine for priming step and the recombinant vaccine for boosting step contain at least one gene of HIV-1 CRFO1AE strain.

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=> e matsuo kazuhiko/au

E1	39	MATSUO KAZUHIRO/AU
E2	120	MATSUO KAZUHIRO/AU
E3	403 -->	MATSUO KAZUHIRO/AU
E4	8	MATSUO KAZUHIRO/AU
E5	10	MATSUO KAZUHIRO/AU
E6	14	MATSUO KAZUHIRO/AU
E7	11	MATSUO KAZUHIRO/AU
E8	18	MATSUO KAZUHIRO/AU
E9	89	MATSUO KAZUHIRO/AU
E10	14	MATSUO KAZUHIRO/AU
E11	8	MATSUO KAZUHIRO/AU
E12	1	MATSUO KAZUHIRO/AU

=> s e3 and bcg

L3 70 "MATSUO KAZUHIRO"/AU AND BCG

=> dup rem l3

PROCESSING COMPLETED FOR L3

L4 47 DUP REM L3 (23 DUPLICATES REMOVED)

=> d bib ab 1-

YOU HAVE REQUESTED DATA FROM 47 ANSWERS - CONTINUE? Y/(N):y

L4 ANSWER 1 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2006:512935 CAPLUS

DN 145:6612

TI A method of prime-boost vaccination for AIDS caused by HIV-1 CRFO1\_AE strain

IN Honda, Mitsuo; Matsuo, Kazuhiko; Hamano, Takaichi; Izumi, Yasuyuki; Promkhatkaew, Duanthanorm; Balachandra, Kruavon; Sutthent, Ruengpung

PA Japan Science and Technology Agency, Japan; Japan as Represented by Director General of National Institute of Infectious Diseases; Departement of Medical Sciences, Ministry of Public Health, Thailand

SO PCT Int. Appl., 39 pp.

CODEN: PIXXD2

DT Patent

LA English

FAN.CNT 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
WO 2006057454	A1	20060601	WO 2005-JP22221	20051125
W:	AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, KE, KG, KM, KN, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, LY, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW			
RW:	AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,			

GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,  
KG, KZ, MD, RU, TJ, TM

JP 2006149234 A 20060615 JP 2004-341283 20041125  
PRAI JP 2004-341283 A 20041125

AB An invention of this application is a method of prime-boost vaccination comprising a priming step by a recombinant BCG vaccine and one or more boosting steps by a recombinant vaccine, wherein both of the recombinant BCG vaccine for priming step and the recombinant vaccine for boosting steps have at least one gene of HIV-1 CRFO1\_AE strain. In said invention, it is a preferred embodiment that both of the recombinant vaccines have at least gag gene of HIV-1 CRFO1-AE strain. The present inventors have found that in the case of using recombinant BCG as a priming antigen in combination with other viral vector-based vaccine as a boosting antigen, quite efficiently enhanced cellular immune response could be induced against HIV-1 CRFO 1\_AE whereupon the present invention has been achieved.

RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 2 OF 47 USPATFULL on STN

AN 2006:247197 USPATFULL

TI Recombination bcg vaccine

IN Honda, Mitsuo, Tokyo, JAPAN

Matsuo, Kazuhiro, Kanagawa, JAPAN

Kanekiyo, Masaru, Tokyo, JAPAN

PI US 2006210586 A1 20060921

AI US 2003-524586 A1 20030813 (10)

WO 2003-JP10303 20030813

20050331 PCT 371 date

PRAI JP 2002-237610 20020816

DT Utility

FS APPLICATION

LREP WENDEROTH, LIND & PONACK, L.L.P., 2033 K STREET N. W., SUITE 800,  
WASHINGTON, DC, 20006-1021, US

CLMN Number of Claims: 2

ECL Exemplary Claim: 1

DRWN 5 Drawing Page(s)

LN.CNT 640

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A recombinant BCG vaccine being transformed with an expression vector that has a polynucleotide encoding a foreign antigenic protein, wherein the polynucleotide is a modified one in which a third position of each codon is substituted with G or C without a change of an amino acid. This recombinant BCG vaccine has an excellent expression rate of antigenic protein and, as a result, capable of inducing a sufficient immune response against target infectious disease, cancer, or the like at the same dose as that of the typical BCG vaccine.

L4 ANSWER 3 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2006:1086167 CAPLUS

DN 145:453348

TI Priming-boosting vaccination with recombinant Mycobacterium bovis bacillus Calmette-Guerin and a nonreplicating vaccinia virus recombinant leads to long-lasting and effective immunity. [Erratum to document cited in CA143:365230]

AU Ami, Yasushi; Izumi, Yasuyuki; Matsuo, Kazuhiro; Someya, Kenji;  
Kanekiyo, Masaru; Horibata, Shigeo; Yoshino, Naoto; Sakai, Koji;  
Shinohara, Katsuaki; Matsumoto, Sohkichi; Yamada, Takeshi; Yamazaki,  
Shudo; Yamamoto, Naoki; Honda, Mitsuo

CS Division of Experimental Animal Research, AIDS Research Center, Division  
of Biosafety Control and Research, National Institute of Infectious  
Diseases, Toyama 1-23-1, Shinjuku-ku, Tokyo, 162-8640, Japan

SO Journal of Virology (2006), 80(20), 10288

CODEN: JOVIAM; ISSN: 0022-538X

PB American Society for Microbiology  
DT Journal  
LA English

AB On page 12873, Table 1, in groups 1, 2 and 3, rBCG was primed at week 3, followed by booster immunization of rDIs at weeks 50 and 54. In the same table, groups 4 and 5, rDIs was primed at weeks 0 and 12, followed by booster immunization of rBCG at week 50. Then all animals were challenged with virulent SHIV KS661c at week 57. On page 12874, Figure 2: The weeks after immunization shown on the x axis in panel A should read: "3, 7, 11, 19, 27, 35, 50, 53, 54, and 56.". The weeks after immunization shown on the x axis in panel B should read: "0, 4, 8, 16, 24, 32, 50, 52, 53, 54, and 56.". Although the patterns and magnitudes of the kinetics were almost the same as the original ones, the standard deviation of the ELISPOT data at the peak response at 54 wk after immunization was 500, which was five times more than originally reported. Spot-forming cells were counted by using a KS ELISPOT system after 35 wk of immunization. Before that, we counted SFCs using an inverted microscope. On page 12878, Acknowledgments, paragraph 1, the last sentence of the paragraph should be deleted. These changes do not alter the conclusions of the article.

L4 ANSWER 4 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN  
AN 2006:1171430 CAPLUS  
DN 146:160902

TI Strategy for development of prophylactic and therapeutic vaccines against HIV

AU Matsuo, Kazuhiro; Yamamoto, Naoki; Honda, Mitsuo  
CS AIDS Research Center, National Institute of Infectious Diseases, Tokyo, Japan

SO Igaku no Ayumi (2006), 218(10), 923-930  
CODEN: IGAYAY; ISSN: 0039-2359

PB Ishiyaku Shuppan  
DT Journal; General Review  
LA Japanese

AB A review discusses development of prophylactic and therapeutic vaccines such as DNA vaccine, BCG vaccine application and antigen peptides vaccine for HIV infection.

L4 ANSWER 5 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN  
DUPLICATE 1

AN 2006:315079 BIOSIS  
DN PREV200600309141

TI Intradermal and oral immunization with recombinant Mycobacterium bovis BCG expressing the simian immunodeficiency virus Gag protein induces long-lasting, antigen-specific immune responses in guinea pigs.

AU Kawahara, Mamoru [Reprint Author]; Matsuo, Kazuhiro; Honda, Mitsuo  
CS Univ Occupat and Environm Hlth, Dept Biochem and Mol Pathophysiol, Sch Med, Yahatanishi Ku, 1-1 Iseigaoka, Kitakyushu, Fukuoka 8078555, Japan  
mamokawa@med.uoeh-u.ac.jp

SO Clinical Immunology (Orlando), (APR 2006) Vol. 119, No. 1, pp. 67-78.  
ISSN: 1521-6616.

DT Article  
LA English

ED Entered STN: 14 Jun 2006  
Last Updated on STN: 14 Jun 2006

AB To develop a new recombinant BCG (rBCG) vaccine, we constructed rBCG that expresses the full-length Gag protein of simian immunodeficiency virus (rBCG-SIVGag) at a level of 0.5 ng/mg after 3 weeks of bacterial cell culture. Intradermal (i.d.) inoculation of guinea pigs with 0.1 mg of rBCG-SIVGag resulted in the induction of delayed-type hypersensitivity (DTH) responses to both purified protein derivative (PPD) of tuberculin and SIV Gag p27 protein; responses that were maintained for the duration of the 50-week study. In contrast, guinea pigs orally vaccinated with 160 mg of the same antigen exhibited a long-lasting DTH response to the SIV

Gag p27 protein, but mounted no response to PPD. Proliferative responses to SIV Gag p27 and PPD antigens were detected in both i.d. and orally immunized animals; however, the levels of PPD-specific responses were significantly higher in guinea pigs immunized by the i.d. than the oral route. A significant increase in the level of PPD- and SIV Gag p27-specific IFN gamma mRNA expression was also detected in both immunization groups receiving rBCG-SIVGag. In addition, both i.d. and oral immunization with rBCG-SIVGag induced PPD- and SIV Gag p27-specific serum IgG responses. Insertion of the SIV gag gene into BCG did not appear to change the ability of rBCG-immunized animals to elicit PPD-specific immune responses. These results indicate that rBCG-SIVGag has the ability to effectively induce long-lasting, cell-mediated and humoral immunity against both viral and bacterial antigens in guinea pigs, suggesting that rBCG-Gag has the potential to elicit immunities specific not only for tuberculosis but also for HIV at human doses. (c) 2005 Elsevier Inc. All rights reserved.

L4 ANSWER 6 OF 47 USPATFULL on STN  
AN 2005:143826 USPATFULL  
TI Radio lan access authentication system  
IN Honda, Mitsuo, Tokyo, JAPAN  
Matsuo, Kazuhiro, Kanagawa, JAPAN  
Izumi, Yasuyuki, Tokyo, JAPAN  
PI US 2005123561 A1 20050609  
AI US 2003-515253 A1 20021120 (10)  
WO 2002-JP12125 20021120  
PRAI JP 2002-145132 20020520  
DT Utility  
FS APPLICATION  
LREP WENDEROTH, LIND & PONACK, L.L.P., 2033 K STREET N. W., SUITE 800,  
WASHINGTON, DC, 20006-1021, US  
CLMN Number of Claims: 7  
ECL Exemplary Claim: 1  
DRWN 14 Drawing Page(s)  
LN.CNT 356  
CAS INDEXING IS AVAILABLE FOR THIS PATENT.  
AB This invention provides a recombinant BCG vaccine transformed by an expression vector having polynucleotide coding for exogenous antigenic protein, characterized in that, the BCG vaccine is used for initial antigen stimulus in immune induction by plural antigen stimulations and also provides a method for the immune induction where the initial antigen stimulus is carried out by the said BCG vaccine and one or more additional antigenic stimulation(s) is/are carried out by non-BCG vaccine expressing the same antigenic protein.

L4 ANSWER 7 OF 47 USPATFULL on STN  
AN 2005:62585 USPATFULL  
TI Recombinant vaccinia virus vaccine  
IN Honda, Mitsuo, Tokyo, JAPAN  
Matsuo, Kazuhiro, Kanagawa, JAPAN  
Ohsu, Takeaki, Kanagawa, JAPAN  
Miyamura, Tatsuo, Tokyo, JAPAN  
Matsuura, Yoshiharu, Osaka, JAPAN  
Ishii, Koji, Tokyo, JAPAN  
Kato, Kenzo, Chiba, JAPAN  
PI US 2005053620 A1 20050310  
AI US 2004-495974 A1 20040903 (10)  
WO 2001-JP10141 20011120  
DT Utility  
FS APPLICATION  
LREP WENDEROTH, LIND & PONACK, L.L.P., 2033 K STREET N. W., SUITE 800,  
WASHINGTON, DC, 20006-1021  
CLMN Number of Claims: 19

ECL Exemplary Claim: 1  
DRWN 7 Drawing Page(s)  
LN.CNT 849

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The invention provides a recombinant vaccinia virus strain DIs that possesses a polynucleotide encoding a foreign antigenic protein in the non-essential gene region of the chromosome DNA and expresses the antigenic protein; and provides a highly-safe, vaccinia virus vaccine containing the recombinant virus strain DIs as the active ingredient. The invention also provides a method of using the vaccinia virus strain DIs as a vector for protein expression.

L4 ANSWER 8 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN  
DUPLICATE 2

AN 2006:8301 BIOSIS

DN PREV200600008996

TI Priming-boosting vaccination with recombinant Mycobacterium bovis bacillus Calmette-Guerin and a nonreplicating vaccinia virus recombinant leads to long-lasting and effective immunity.

AU Ami, Yasushi; Izumi, Yasuyuki; Matsuo, Kazuhiro; Someya, Kenji; Kanekiyo, Masaru; Horibata, Shigeo; Yoshino, Naoto; Sakai, Koji; Shinohara, Katsuaki; Matsumoto, Sohkiichi; Yamada, Takeshi; Yamazaki, Shudo; Yamamoto, Naoki; Honda, Mitsuo [Reprint Author]

CS Natl Inst Infect Dis, Ctr AIDS Res, Shinjuku Ku, Toyama 1-23-1, Tokyo 1628640, Japan  
mhonda@nih.go.jp

SO Journal of Virology, (OCT 2005) Vol. 79, No. 20, pp. 12871-12879.  
CODEN: JOVIAM. ISSN: 0022-538X.

DT Article

LA English

ED Entered STN: 14 Dec 2005

Last Updated on STN: 14 Dec 2005

AB Virus-specific T-cell responses can limit immunodeficiency virus type 1 (HIV-1) transmission and prevent disease progression and so could serve as the basis for an affordable, safe, and effective vaccine in humans. To assess their potential for a vaccine, we used Mycobacterium bovis bacillus Calmette-Guerin (BCG)-Tokyo and a replication-deficient vaccinia virus strain (DIs) as vectors to express full-length gag from simian immunodeficiency viruses (SIVs) (rBCG-SIVgag and rDisSIVgag). Cynomolgus macaques were vaccinated with either rBCG-SIVgag dermally as a single modality or in combination with rDisSIVgag intravenously. When cynomolgus macaques were primed with rBCG-SIVgag and then boosted with rDisSIVgag, high levels of gamma interferon (IFN-gamma) spot-forming cells specific for SIV Gag were induced. This combination regimen elicited effective protective immunity against mucosal challenge with pathogenic simian-human immunodeficiency virus for the 1 year the macaques were under observation. Antigen-specific intracellular IFN-gamma activity was similarly induced in each of the macaques with the priming-boosting regimen. Other groups receiving the opposite combination or the single-modality vaccines were not effectively protected. These results suggest that a recombinant M. bovis BCG-based vector may have potential as an HIV/AIDS vaccine when administered in combination with a replication-deficient vaccinia virus DIs vector in a priming-boosting strategy.

L4 ANSWER 9 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN  
DUPLICATE 3

AN 2005:399376 BIOSIS

DN PREV200510190449

TI Mycobacterial codon optimization enhances antigen expression and virus-specific immune responses in recombinant Mycobacterium bovis bacille Calmette-Guerin expressing human immunodeficiency virus type 1 Gag.

AU Kanekiyo, Masaru; Matsuo, Kazuhiro; Hamatake, Makiko; Hamano, Takaichi; Ohsu, Takeaki; Matsumoto, Sohkiichi; Yamada, Takeshi; Yamazaki,

Shudo; Hasegawa, Atsuhiko; Yamamoto, Naoki; Honda, Mitsuo [Reprint Author]  
CS Natl Inst Infect Dis, AIDS Res Ctr, Shinjuku Ku, 1-23-1 Toyama, Tokyo  
1628640, Japan  
mhonda@nih.go.jp  
SO Journal of Virology, (JUL 2005) Vol. 79, No. 14, pp. 8716-8723.  
CODEN: JOVIAM. ISSN: 0022-538X.

DT Article

LA English

ED Entered STN: 5 Oct 2005

Last Updated on STN: 5 Oct 2005

AB Although its potential for vaccine development is already known, the introduction of recombinant human immunodeficiency virus (HIV) genes to *Mycobacterium bovis* bacille Calmette-Guerin (BCG) has thus far elicited only limited responses. In order to improve the expression levels, we optimized the codon usage of the HIV type 1 (HIV-1) p24 antigen gene of gag (p24 gag) and established a codon-optimized recombinant BCG (rBCG)-p24 Gag which expressed a 40-fold-higher level of p24 Gag than did that of nonoptimized rBCG-p24 Gag. Inoculation of mice with the codon-optimized rBCG-p24 Gag elicited effective immunity, as evidenced by virus-specific lymphocyte proliferation, gamma interferon ELISPOT cell induction, and antibody production. In contrast, inoculation of animals with the nonoptimized rBCG-p24 Gag induced only low levels of immune responses. Furthermore, a dose as small as 0.01 mg of the codon-optimized rBCG per animal proved capable of eliciting immune responses, suggesting that even low doses of a codon-optimized rBCG-based vaccine could effectively elicit HIV-1-specific immune responses.

L4 ANSWER 10 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 4

AN 2005:700913 CAPLUS

DN 143:210149

TI Ag85B of *Mycobacteria* Elicits Effective CTL Responses through Activation of Robust Th1 Immunity as a Novel Adjuvant in DNA Vaccine

AU Takamura, Shiki; Matsuo, Kazuhiro; Takebe, Yutaka; Yasutomi, Yasuhiro

CS Japanese Foundation for AIDS Prevention, Tokyo, Japan

SO Journal of Immunology (2005), 175(4), 2541-2547

CODEN: JOIMA3; ISSN: 0022-1767

PB American Association of Immunologists

DT Journal

LA English

AB CD4+ T cells play a crucial role in CTL generation in a DNA vaccination strategy. Several studies have demonstrated the requirement of CD4+ T cells for the induction of a sufficient immune response by coadministering DNAs. In the present study the authors investigated the effectiveness of Ag85B of *mycobacteria*, which is known to be one of the immunogenic proteins for Th1 development, as an adjuvant of a DNA vaccine. HIV gp120 DNA vaccine mixed with Ag85B DNA as an adjuvant induced HIV gp120-specific Th1 responses, as shown by delayed-type hypersensitivity, cytokine secretion, and increasing HIV-specific CTL responses. Moreover, these responses were enhanced in mice primed with *Mycobacterium bovis* bacillus Calmette-Guerin before immunization of HIV DNA vaccine mixed with Ag85B DNA. Furthermore, these immunized mice showed substantial reduction of HIV gp120-expressing recombinant vaccinia virus titers compared with the titers in other exptl. mice after recombinant vaccinia virus challenge. Because most humans have been sensitized by spontaneous infection or by vaccination with *mycobacteria*, these findings indicate that Ag85B is a promising adjuvant for enhancing CTL responses in a DNA vaccination strategy.

RE.CNT 55 THERE ARE 55 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 11 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 5

AN 2005:114055 CAPLUS

DN 142:334776

TI Vaccination of rhesus macaques with recombinant Mycobacterium bovis  
 bacillus Calmette-Guerin Env V3 elicits neutralizing antibody-mediated  
 protection against simian-human immunodeficiency virus with a homologous  
 but not a heterologous V3 motif

AU Someya, Kenji; Cecilia, Dayaraj; Ami, Yasushi; Nakasone, Tadashi;  
 Matsuo, Kazuhiro; Burda, Sherri; Yamamoto, Hiroshi; Yoshino,  
 Naoto; Kaizu, Masahiko; Ando, Shuji; Okuda, Kenji; Zolla-Pazner, Susan;  
 Yamazaki, Shudo; Yamamoto, Naoki; Honda, Mitsuo

CS AIDS Research Center, National Institute of Infectious Diseases, Tokyo,  
 162-8640, Japan

SO Journal of Virology (2005), 79(3), 1452-1462  
 CODEN: JOVIAM; ISSN: 0022-538X

PB American Society for Microbiology

DT Journal

LA English

AB Although the correlates of vaccine-induced protection against human  
 immunodeficiency virus type 1 (HIV-1) are not fully known, it is presumed  
 that neutralizing antibodies (NAb) play a role in controlling virus  
 infection. In this study, we examined immune responses elicited in rhesus  
 macaques following vaccination with recombinant Mycobacterium bovis  
 bacillus Calmette-Guerin expressing an HIV-1 Env V3 antigen (rBCG Env V3).  
 We also determined the effect of vaccination on protection against challenge  
 with either a simian-human immunodeficiency virus (SHIV-MN) or a highly  
 pathogenic SHIV strain (SHIV-89.6PD). Immunization with rBCG Env V3  
 elicited significant levels of NAb for the 24 wk tested that were  
 predominantly HIV-1 type specific. Sera from the immunized macaques  
 neutralized primary HIV-1 isolates in vitro, including HIV-1BZ167/X4,  
 HIV-1SF2/X4, HIV-1C12/X4, and, to a lesser extent, HIV-1MNp/X4, all of  
 which contain a V3 sequence homologous to that of rBCG Env V3. In  
 contrast, neutralization was not observed against HIV-1SF33/X4, which has a  
 heterologous V3 sequence, nor was it found against primary HIV-1 R5  
 isolates from either clade A or B. Furthermore, the viral load in the  
 vaccinated macaques was significantly reduced following low-dose challenge  
 with SHIV-MN, and early plasma viremia was markedly decreased after  
 high-dose SHIV-MN challenge. In contrast, replication of pathogenic  
 SHIV-89.6PD was not affected by vaccination in any of the macaques. Thus,  
 we have shown that immunization with an rBCG Env V3 vaccine elicits a  
 strong, type-specific V3 NAb response in rhesus macaques. While this  
 response was not sufficient to provide protection against a pathogenic  
 SHIV challenge, it was able to significantly reduce the viral load in  
 macaques following challenge with a nonpathogenic SHIV. These  
 observations suggest that rBCG vectors have the potential to deliver an  
 appropriate virus immunogen for desirable immune elicitation.

RE.CNT 54 THERE ARE 54 CITED REFERENCES AVAILABLE FOR THIS RECORD  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 12 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2005:1019515 CAPLUS

DN 144:86200

TI BCG as a vaccine vector

AU Matsuo, Kazuhiro

CS First Research Group, AIDS Research Center, National Institute of  
 Infectious Diseases, Tokyo, 162-8640, Japan

SO Rinsho Men'eki (2005), 43(6), 659-664  
 CODEN: RNMKAU; ISSN: 0386-9695

PB Kagaku Hyoronsha

DT Journal; General Review

LA Japanese

AB A review discusses application of BCG as vectors for vaccines  
 such as AIDS vaccines.

L4 ANSWER 13 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
 STN DUPLICATE 6

AN 2005:338939 BIOSIS



DN PREV200510120020  
 TI Specific immune response and pathological findings in BALB/c mice inoculated with recombinant BCG expressing HIV-1 antigen.  
 AU Wiriyarat, Witthawat; Sukpanichnant, Sanya; Sittisombut, Nopporn; Balachandra, Kruavon; Promkhatkaew, Duanthanorm; Butraporn, Raywadee; Sutthent, Ruengpung; Boonlong, Jotika; Matsuo, Kazuhiro; Honda, Mitsuo; Warachit, Paijit; Puthavathana, Pilaipan [Reprint Author]  
 CS Mahidol Univ, Siriraj Hosp, Fac Med, Dept Microbiol, Bangkok 10700, Thailand  
 siput@mahidol.ac.th  
 SO Asian Pacific Journal of Allergy and Immunology, (MAR 2005) Vol. 23, No. 1, pp. 41-51.  
 ISSN: 0125-877X.  
 DT Article  
 LA English  
 ED Entered STN: 31 Aug 2005  
 Last Updated on STN: 31 Aug 2005  
 AB Recombinant BCGs (rBCGs) containing extrachromosomal plasmids with different HIV-1 insert sequences: nef, env (V3J1 and E9Q), gag p17 or whole gag p55 were evaluated for their immunogenicity, safety and persistent infection in BALB/c mice. Animal injected with, rBCG-pIJKV3J1, rBCG-pSO gag p17 or rBCG-pSO gag p55 could elicit lymphocyte proliferation as tested by specific HIV-1 peptides or protein antigen. Inoculation with various concentration of rBCG-pSO gag p55 generated satisfactory specific lymphocyte proliferation in dose escalation trials. The rBCG-pSO gag p55 recovered from spleen tissues at different time interval post-inoculation could express the HIV protein as determined by ELISA p24 antigen detection kit. This result indicated that the extrachromosomal plasmid was stable and capable to express Gag protein. It was also demonstrated that rBCGs did not cause serious pathological change in the inoculated animals. The present study suggested the role of BCG as a potential vehicle for using in HIV vaccine development.

L4 ANSWER 14 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN  
 AN 2004:162604 CAPLUS  
 DN 140:216160  
 TI Recombinant BCG vaccines against infection, cancer and other diseases  
 IN Honda, Mitsuo; Matsuo, Kazuhiro; Kanekiyo, Masaru  
 PA Japan Science and Technology Corporation, Japan; Japan as Represented by Director General of National Institute of Infectious Diseases  
 SO PCT Int. Appl., 31 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2004016280	A1	20040226	WO 2003-JP10303	20030813
	W: IN, JP, US				
	RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR				
	EP 1535627	A1	20050601	EP 2003-788098	20030813
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, SK				
	US 2006210586	A1	20060921	US 2005-524586	20050331
PRAI	JP 2002-237610	A	20020816		
	WO 2003-JP10303	W	20030813		

AB A recombinant BCG vaccine obtained by transformation with an expression vector carrying a polynucleotide coding for an ecdemic antigenic protein, which recombinant BCG vaccine consists of a modified type polynucleotide comprising a polynucleotide having the third base of each of the codons thereof substituted with G or C without changing of the type of amino acid. This recombinant BCG

vaccine excels in the amount of antigenic protein expressed, so that even with the same dosage as employed for conventional BCG vaccines, the recombinant BCG vaccine can induce satisfactory immune response to target infectious diseases, cancer, etc. In example, recombinant BCG comprising hsp60 gene and HIV-1 gag p24 gene was prepared for use as vaccine.

RE.CNT 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 15 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2003:931202 CAPLUS

DN 139:394884

TI BCG vector and non-BCG vector encoding antigenic protein for use as vaccine and booster vaccine against infection and cancer

IN Honda, Mitsuo; Matsuo, Kazuhiro; Izumi, Yasuyuki

PA Japan Science and Technology Corporation, Japan; Japan as Represented by Director General of National Institute of Infectious Diseases

SO PCT Int. Appl., 35 pp.

CODEN: PIXXD2

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2003097087	A1	20031127	WO 2002-JP12125	20021120
	W: CA, JP, US				
	CA 2494359	A1	20031127	CA 2002-2494359	20021120
	US 2005123561	A1	20050609	US 2003-515253	20021120
	IN 2003DE00709	A	20050311	IN 2003-DE709	20030520
PRAI	JP 2002-145132	A	20020520		
	WO 2002-JP12125	W	20021120		

AB A recombinant BCG vaccine transformed by an expression vector which carries a polynucleotide encoding a foreign antigenic protein characterized by being usable for priming in immune induction via twice or more antigenic stimulations; and an immune induction method characterized by comprising priming with the use of the above BCG vaccine and boosting once or more with the use of a non-BCG vaccine expressing the same antigenic protein. Recombinant BCG (rBCG-SIVgag) and recombinant vaccinia Dis (rDis-SIVgaga) encoding SIV gag gene were prepared and tested.

RE.CNT 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD  
ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 16 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN  
DUPLICATE 7

AN 2003:346612 BIOSIS

DN PREV200300346612

TI Immunization with recombinant Calmette-Guerin bacillus (BCG)-hepatitis C virus (HCV) elicits HCV-specific cytotoxic T lymphocytes in mice.

AU Uno-Furuta, Satori; Matsuo, Kazuhiro; Tamaki, Shigenori; Takamura, Shiki; Kamei, Akira; Kuromatsu, Isao; Kaito, Masahiko; Matsuura, Yoshiharu; Miyamura, Tatsuo; Adachi, Yukihiko; Yasutomi, Yasuhiro [Reprint Author]

CS Department of Bioregulation, Mie University School of Medicine, 2-174 Edobashi, Tsu, Mie, 514-8507, Japan  
yasutomi@doc.medic.mie-u.ac.jp

SO Vaccine, (4 July 2003) Vol. 21, No. 23, pp. 3149-3156. print.  
ISSN: 0264-410X (ISSN print).

DT Article

LA English

ED Entered STN: 30 Jul 2003

Last Updated on STN: 30 Jul 2003

AB Since virus-specific cytotoxic T lymphocytes (CTLs) play a critical role in preventing the spread of hepatitis C virus (HCV), an effective HCV vaccine should be capable of eliciting HCV-specific CTLs. In the present study, we assessed the capability of a novel recombinant vaccine using an attenuated tuberculosis bacillus, Calmette-Guerin bacillus (BCG), as a vaccine vehicle to elicit HCV-specific CTLs. BCG was engineered to express the CTL epitope of HCV-non-structure protein 5a (NS5a) as a chimeric protein with alpha antigen of mycobacteria. Immunization with this recombinant BCG elicited major histocompatibility complex class I-restricted CD8+ HCV-NS5a-specific CTLs in mice. Immunized mice showed a substantial reduction in the vaccinia virus titer compared with control mice when the immunized mice were challenged with a recombinant vaccinia virus expressing HCV-NS5a genes. These findings provide evidences for the possibility of BCG as a vaccine vector and its continued exploration as a vehicle for eliciting HCV-specific immunity.

L4 ANSWER 17 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2003:909231 CAPLUS

DN 140:31200

TI New trend in AIDS vaccine research

AU Matsuo, Kazuhiro

CS AIDS Research Center, National Institute of Infectious Diseases, Japan

SO Kagaku to Seibutsu (2003), 41(11), 731-737

CODEN: KASEAA; ISSN: 0453-073X

PB Gakkai Shuppan Senta

DT Journal; General Review

LA Japanese

AB A review on infection with HIV-1, symptoms of AIDS, shifting from neutralizing antibody-inducing vaccines to cell-mediated immunity-inducing vaccines, vaccines currently being clin. tested, and immune induction by combination of recombinant BCG and recombinant vaccinia virus.

L4 ANSWER 18 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 2002:169547 CAPLUS

DN 136:231223

TI Inducer of Th1-type cytokine production and it's application as vaccine for treating atopic diseases

IN Yasutomi, Yasuhiro; Mizutani, Hitoshi; Matsuo, Kazuhiro

PA Japan BCG Laboratory, Japan

SO Jpn. Kokai Tokkyo Koho, 4 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	JP 2002068990	A	20020308	JP 2000-261016	20000830
PRAI	JP 2000-261016		20000830		

AB Provided is an agent capable of switching cytokine profiles from Th2 to Th1, or inducing Th1 cytokine production. The Th1 cytokine inducer is heat-killed/ultrasound-disrupted BCG or components derived from it. The inducer can be used as a vaccine to treat atopic diseases such as atopic dermatitis and asthma.

L4 ANSWER 19 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 8

AN 2003:19497 BIOSIS

DN PREV200300019497

TI Combined intrarectal/intradermal inoculation of recombinant Mycobacterium bovis bacillus Calmette-Guerin (BCG) induces enhanced immune responses against the inserted HIV-1 V3 antigen.

AU Kawahara, Mamoru; Matsuo, Kazuhiro; Nakasone, Tadashi; Hiroi, Takachika; Kiyono, Hiroshi; Matsumoto, Sohkiichi; Yamada, Takeshi;

Yamamoto, Naoki; Honda, Mitsuo [Reprint Author]  
 CS AIDS Research Center, National Institute of Infectious Diseases, 1-23-1  
 Toyama, Shinjuku-ku, Tokyo, 162-8640, Japan  
 mhonda@nih.go.jp  
 SO Vaccine, (13 December 2002) Vol. 21, No. 3-4, pp. 158-166. print.  
 ISSN: 0264-410X (ISSN print).  
 DT Article  
 LA English  
 ED Entered STN: 1 Jan 2003  
 Last Updated on STN: 1 Jan 2003  
 AB The development of a successful recombinant Mycobacterium bovis bacillus  
 Calmette-Guerin (rBCG) vector-based vaccine for human immunodeficiency  
 virus type 1 (HIV-1) requires the induction of high levels of  
 HIV-1-specific immunity while at the same time maintaining immunity to  
 tuberculosis. To examine a combined vaccination strategy for enhancement  
 of immune responses specific for HIV-1, guinea pigs were inoculated with  
 either a single or combination intradermal (i.d.), intrarectal (i.r.) and  
 intranasal (i.n.) administration of rBCG-pSOV3J1 which secretes a chimeric  
 protein of HIV-1 V3J1 peptide and alpha-antigen. Significant level of  
 delayed-type hypersensitivity to both V3J1 peptide and tuberculin was  
 induced in guinea pigs inoculated with human doses of rBCG-pSOV3J1 by a  
 combination of intrarectal and intradermal routes. Guinea pigs inoculated  
 by combined routes also had significantly higher titers of HIV-1-specific  
 serum IgG and IgA compared with those animals immunized only  
 intrarectally, which led to the enhanced neutralization activity against  
 HIV-1MN. In addition, the induction of high levels of IFNgamma and  
 interleukin-2 (IL-2) mRNA in PBMC, splenocytes, and intraepithelial  
 lymphocytes from the immunized animals was detected until at least 110  
 weeks post-inoculation. These results suggest that enhanced immune  
 responses specific for HIV-1 are efficiently induced by combined  
 intrarectal and intradermal immunization with rBCG-HIV, and  
 antigen-specific Th1-type memory cells are maintained for more than 2  
 years in the immunized animals. Thus, inoculation with rBCG-HIV by  
 combined routes represents an effective vaccination strategy to elicit  
 high levels of HIV-1-specific immune responses.

L4 ANSWER 20 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
 STN  
 AN 2002:634285 BIOSIS  
 DN PREV200200634285  
 TI Protective efficacy of vaccine candidates against guinea pig pulmonary  
 tuberculosis.  
 AU Yamamoto, Saburo; Yamamoto, Toshiko; Nojima, Yasuhiro; Umemori, Kiyoko;  
 Matsuo, Kazuhiro; Nomaguchi, Hiroko; Sato, Yukio; Yamada, Takeshi;  
 Ohara, Naoya; Matsumoto, Sohkiichi; Phalen, Susan; McMurray, David N.  
 SO Tuberculosis (Edinburgh), (2002) Vol. 82, No. 2-3, pp. 128. print.  
 Meeting Info.: 36th Annual Research Conference of the US-Japan Cooperative  
 Medical Science Program Tuberculosis and Leprosy Panel. Louisiana, USA.  
 July 15-17, 2001.  
 ISSN: 1472-9792.  
 DT Conference; (Meeting)  
 Conference; Abstract; (Meeting Abstract)  
 LA English  
 ED Entered STN: 12 Dec 2002  
 Last Updated on STN: 12 Dec 2002

L4 ANSWER 21 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN  
 AN 2002:611508 CAPLUS  
 DN 138:50341  
 TI Discovery of immunostimulatory CpG-DNA and its application to tuberculosis  
 vaccine development  
 AU Yamamoto, Saburo; Yamamoto, Toshiko; Nojima, Yasuhiro; Umemori, Kiyoko;  
 Phalen, Susan; McMurray, David N.; Kuramoto, Etsuro; Iho, Sumiko; Takauji,  
 Rumiko; Sato, Yukio; Yamada, Takeshi; Ohara, Naoya; Matsumoto, Sohkiichi;

Goto, Yoshitaka; Matsuo, Kazuhiro; Tokunaga, Tohru  
 CS National Institute of Infectious Diseases, Musashimurayama, Tokyo,  
 208-0011, Japan  
 SO Japanese Journal of Infectious Diseases (2002), 55(2), 37-44  
 CODEN: JJIDFE; ISSN: 1344-6304  
 PB National Institute of Infectious Diseases  
 DT Journal; General Review  
 LA English  
 AB A review. DNA containing an unmethylated CpG motif has a potent immunostimulatory effect on the vertebrate immune system. Because such CpG motifs are relatively common in bacterial DNA, but rare in mammalian animal and plant DNA, they may be an evolutionary adaptation augmenting innate immunity, most likely in response to pathogens that replicate within the host cells, such as viruses and intracellular bacteria. Microbial infection induces innate immunity by triggering pattern-recognition systems. The infected cells produce proinflammatory cytokines that directly combat microbial invaders and express costimulating surface mols., which develop adaptive immunity by inducing distinct T cell differentiation. Bacterial DNA with unmethylated CpG-DNA stimulates vertebrate immature immune cells to induce maturation and to produce TNF- $\alpha$  as well as Th1-type cytokines, IL-12 and IFN- $\gamma$ . Therefore, CpG-DNA functions as an adjuvant for regulating the initiation of Th1 differentiation. The roles of immunostimulatory CpG motifs in DNA vaccine developments and in therapeutic applications have been discussed.

RE.CNT 58 THERE ARE 58 CITED REFERENCES AVAILABLE FOR THIS RECORD  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 22 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
 STN DUPLICATE 9  
 AN 2002:89940 BIOSIS  
 DN PREV200200089940  
 TI Cross-clade neutralizing antibody production against human immunodeficiency virus type 1 clade E and B' strains by recombinant Mycobacterium bovis BCG-based candidate vaccine.  
 AU Chujoh, Yoshitomo; Matsuo, Kazuhiro; Yoshizaki, Hitomi; Nakasatomi, Tetsuya; Someya, Kenji; Okamoto, Yukari; Naganawa, Satoshi; Haga, Shinji; Yoshikura, Hiroshi; Yamazaki, Akihiro; Yamazaki, Shudo; Honda, Mitsuo [Reprint author]  
 CS AIDS Research Center, National Institute of Infectious Diseases, Toyama 1-23-1, Shinjuku-ku, Tokyo, 162-8640, Japan  
 mhonda@nih.go.jp  
 SO Vaccine, (12 December, 2001) Vol. 20, No. 5-6, pp. 797-804. print.  
 CODEN: VACCDE. ISSN: 0264-410X.  
 DT Article  
 LA English  
 ED Entered STN: 24 Jan 2002  
 Last Updated on STN: 25 Feb 2002  
 AB The recombinant Mycobacterium bovis BCG (rBCG) vector-based vaccine secreting the V3 principal neutralizing epitope of human immunodeficiency virus type 1 (HIV-1) Japanese strain was reported to induce both humoral and cellular immune responses effectively (Proc. Natl. Acad. Sci. USA. 92 (1995) 10693). The antigen-secreting rBCG system was applied to the V3 epitope of clade E HIV-1 in this study. The V3 sequence of 19 amino acids (aa) and 15aa fused with mycobacterial alpha-antigen was not secreted while 12aa and 11aa sequences were successfully secreted from BCG cells. Serum IgG from guinea pig which was immunized with 12aa epitope-secreting recombinant BCG neutralized the WHO reference strain as well as primary field isolates of clade E virus. The serum IgG could also neutralize Thai B (clade B') strains which possessed a conserved GPGQ motif in their V3 sequences. These data suggest that the rBCG construct secreting the 12aa epitope is implicated in the development of a prophylactic vaccine in Thailand in which both clade E and B' viruses are prevalent.

L4 ANSWER 23 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
 STN DUPLICATE 10  
 AN 2002:479818 BIOSIS  
 DN PREV200200479818  
 TI Induction of effective antitumor immune responses in a mouse bladder tumor  
 model by using DNA of an alpha antigen from mycobacteria.  
 AU Kuromatsu, Isao; Matsuo, Kazuhiro; Takamura, Shiki; Kim, Gisen;  
 Takebe, Yutaka; Kawamura, Juichi; Yasutomi, Yasuhiro [Reprint author]  
 CS Department of Bioregulation, Mie University School of Medicine, 2-174  
 Edobashi, Tsu, Mie, 514-8507, Japan  
 yasutomi@doc.medic.mie-u.ac.jp  
 SO Cancer Gene Therapy, (July, 2001) Vol. 8, No. 7, pp. 483-490. print.  
 ISSN: 0929-1903.  
 DT Article  
 LA English  
 ED Entered STN: 11 Sep 2002  
 Last Updated on STN: 11 Sep 2002  
 AB One of the main objectives of cancer immunotherapy is the activation and  
 increase in number of antitumor effector cells. Recently, genetically  
 modified tumor cell vaccines have been proposed for elicitation of  
 antitumor effector cells. Native alpha antigen (alpha Ag) (also known as  
 MPT59 and antigen 85B) of mycobacteria, which cross-reacts among  
 mycobacteria species, may play an important biological role in  
 host-pathogen interaction because it elicits various helper T-cell type 1  
 immune responses. To assess the induction of antitumor immune responses  
 by alpha Ag, mouse tumor cell lines transfected with cDNA of alpha Ag from  
 Mycobacterium kansasii were established, and the possibility of producing  
 a tumor cell vaccine for induction of antitumor effects was explored.  
 Transfection of tumor cell lines with an alpha Ag gene lead to primary  
 tumor rejection and the establishment of protective immunity to  
 nontransfected original tumor cell lines in Mycobacterium bovis bacillus  
 Calmette-Guerin (BCG)-primed and unprimed mice. Mice immunized  
 with tumor cell lines transfected with the alpha Ag gene showed  
 delayed-type hypersensitivity responses in vivo and proliferative  
 responses together with induction of interferon-gamma of spleen cells  
 against nontransfected wild-type tumor cell lines in in vitro experiments.  
 Moreover, immunization of mice with alpha Ag-expressing tumor cells  
 elicited tumor-specific and cytotoxic T lymphocyte (CTL) epitope  
 peptide-specific CD8+ CTLs. The results of this study provided evidence  
 of the potential usefulness of alpha Ag in tumor cell vaccines.

L4 ANSWER 24 OF 47 USPATFULL on STN  
 AN 2000:7197 USPATFULL  
 TI Mycobacterial secretory expression vectors and transformants  
 IN Yamada, Takeshi, Nagasaki-ken, Japan  
 Matsuo, Kazuhiro, Kaswaski, Japan  
 Yamaguchi, Ryuji, Kawasaki, Japan  
 Yamazaki, Akihiro, Kawasaki, Japan  
 PA Ajinomoto, Co., Inc., Tokyo, Japan (non-U.S. corporation)  
 Takeshi Yamada, Nagasaki-ken, Japan (non-U.S. corporation)  
 PI US 6015696 20000118  
 AI US 1994-193899 19940209 (8)  
 RLI Continuation of Ser. No. US 1990-531448, filed on 31 May 1990, now  
 abandoned  
 PRAI JP 1989-135855 19890531  
 JP 1990-64310 19900316  
 DT Utility  
 FS Granted  
 EXNAM Primary Examiner: Railey, II, Johnny F.  
 LREP Oblon, Spivak, McClelland, Maier & Neustadt, P.C.  
 CLMN Number of Claims: 9  
 ECL Exemplary Claim: 1  
 DRWN 6 Drawing Figure(s); 6 Drawing Page(s)  
 LN.CNT 737

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A secretory vector expressed in mycobacteria comprising a promoter, a signal sequence having ligated therewith a DNA nucleotide sequence encoding heterologous polypeptide and replicator region capable of replication in mycobacteria is disclosed. A transformant that has been transformed with the secretory vector, as well as a vaccine comprising the transformant is provided.

L4 ANSWER 25 OF 47 USPATFULL on STN  
AN 1999:36712 USPATFULL  
TI Anti-AIDS secretory recombinant BCG vaccine  
IN Matsuo, Kazuhiro, Kawasaki, Japan  
Chujo, Yoshitomo, Kawasaki, Japan  
Yamazaki, Akihiro, Kawasaki, Japan  
Honda, Mitsuo, Mitaka, Japan  
Yamazaki, Shudo, Higashiyamato, Japan  
Tasaka, Hiromichi, Kure, Japan  
PA Ajinomoto Co., Inc., Tokyo, Japan (non-U.S. corporation)  
Japan as represented by Director General of Agency of National Institute of Health, Tokyo, Japan (non-U.S. government)  
PI US 5885580 19990323  
AI US 1997-972089 19971117 (8)  
RLI Division of Ser. No. US 1996-619512, filed on 29 Mar 1996, now abandoned  
PRAI JP 1994-178462 19940729  
DT Utility  
FS Granted  
EXNAM Primary Examiner: Stucker, Jeffrey; Assistant Examiner: Park, Hankyel T.  
LREP Oblon, Spivak, McClelland, Maier & Neustadt, P.C.  
CLMN Number of Claims: 26  
ECL Exemplary Claim: 1  
DRWN 15 Drawing Figure(s); 11 Drawing Page(s)  
LN.CNT 1556

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A vaccine containing Mycobacterium bovis BCG which secretes a fusion protein to be obtained by inserting a foreign antigen peptide into the molecular surface of a secretory protein, a carrier, having a signal peptide. BCG constituting the present invention secretes a fusion protein to be obtained by inserting a foreign antigen peptide into the molecular surface of an  $\alpha$ -antigen derived from mycobacteria. Said fusion protein has significantly increased antigenicity and immunogenicity. Therefore, when it is inoculated into animals, it is efficiently recognized by B cells which recognize said antigen, thereby effectively inducing the production of an antibody to said antigen. When said BCG itself is inoculated into animals, it continuously secretes said fusion protein in the bodies of the animals while continuously propagating therein. Therefore, said BCG is an extremely useful vaccine.

L4 ANSWER 26 OF 47 USPATFULL on STN  
AN 1999:4042 USPATFULL  
TI Anti-acids secretory recombinant BCG vaccine  
IN Matsuo, Kazuhiro, Kawasaki, Japan  
Chujo, Yoshitomo, Kawasaki, Japan  
Yamazaki, Akihiro, Kawasaki, Japan  
Honda, Mitsuo, Mitaka, Japan  
Yamazaki, Shudo, Higashiyamato, Japan  
Tasaka, Hiromichi, Kure, Japan  
PA Ajinomoto Co., Inc., Tokyo, Japan (non-U.S. corporation)  
Japan as represented by Director General of Agency of National Institute of Health, Tokyo, Japan (non-U.S. corporation)  
PI US 5858369 19990112  
AI US 1997-975699 19971121 (8)  
RLI Continuation of Ser. No. US 1996-619512, filed on 29 Mar 1996, now abandoned

PRAI JP 1994-178462 19940729  
 DT Utility  
 FS Granted  
 EXNAM Primary Examiner: Stucker, Jeffrey; Assistant Examiner: Park, Hankyel T.  
 LREP Oblon, Spivak, McClelland, Maier & Neustadt, P.C.  
 CLMN Number of Claims: 16  
 ECL Exemplary Claim: 1  
 DRWN 15 Drawing Figure(s); 11 Drawing Page(s)  
 LN.CNT 1514

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A vaccine containing Mycobacterium bovis BCG which secretes a fusion protein to be obtained by inserting a foreign antigen peptide into the molecular surface of a secretory protein, a carrier, having a signal peptide. BCG constituting the present invention secretes a fusion protein to be obtained by inserting a foreign antigen peptide into the molecular surface of an  $\alpha$ -antigen derived from mycobacteria. Said fusion protein has significantly increased antigenicity and immunogenicity. Therefore, when it is inoculated into animals, it is efficiently recognized by B cells which recognize said antigen, thereby effectively inducing the production of an antibody to said antigen. When said BCG itself is inoculated into animals, it continuously secretes said fusion protein in the bodies of the animals while continuously propagating therein. Therefore, said BCG is an extremely useful vaccine.

L4 ANSWER 27 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN  
 AN 1996:340573 CAPLUS  
 DN 125:8464  
 TI Anti-AIDS secretory recombinant BCG vaccine containing  $\alpha$ -antigen of acid-fast bacterium to enhance antigenicity  
 IN Matsuo, Kazuhiro; Chujo, Yoshitomo; Yamazaki, Akihiro; Honda, Mitsuo; Yamazaki, Shudo; Tasaka, Hiromichi  
 PA Ajinomoto Co., Inc., Japan; Japan, Agency of National Institute of Health  
 SO PCT Int. Appl., 56 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA Japanese  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 9604009	A1	19960215	WO 1995-JP1515	19950731
	W: CN, JP, KR, US				
	RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE				
	CN 1136280	A	19961120	CN 1995-190959	19950731
	EP 745386	A1	19961204	EP 1995-926523	19950731
	EP 745386	B1	20040204		
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE				
	AT 258986	T	20040215	AT 1995-926523	19950731
	US 5885580	A	19990323	US 1997-972089	19971117
	US 5858369	A	19990112	US 1997-975699	19971121
PRAI	JP 1994-178462	A	19940729		
	WO 1995-JP1515	W	19950731		
	US 1996-619512	B3	19960329		

AB A vaccine containing Mycobacterium bovis BCG which secretes a fused protein obtained by inserting a foreign antigen peptide to the surface of a mol. of a secreted protein having a signal peptide as a carrier. The BCG secretes a fused protein obtained by inserting a foreign antigen peptide to the surface of a mol. of an  $\alpha$ -antigen originating in an acid-fast bacterium. The fused protein has extremely enhanced antigenicity and can efficiently induce the production of an antibody against the  $\alpha$ -antigen in animals. When the BCG can serve as a live vaccine; it continues to grow and secrete the fused protein in the animal body. Preparation of recombinant secretion vectors encoding fusion protein containing  $\alpha$ -antigen and the epitope of V3 surface antigen of



HIV-1 was shown and their antigenicity were demonstrated in vivo. The antibodies thus produced also cross-react to V3 epitopes of types Thai-A and Thai-B.

- L4 ANSWER 28 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
STN DUPLICATE 11
- AN 1996:21760 BIOSIS
- DN PREV199698593895
- TI Protective immune responses induced by secretion of a chimeric soluble protein from a recombinant *Mycobacterium bovis* bacillus Calmette-Guerin vector candidate vaccine for human immunodeficiency virus type 1 in small animals.
- AU Honda, Mitsuo [Reprint author]; Matsuo, Kazuhiro; Nakasone, Tadashi; Okamoto, Yukari; Yoshizaki, Hitomi; Kitamura, Katsuhiko; Sugiura, Wataru; Watanabe, Kuhomi; Fukushima, Yoshiko; Haga, Shinji; Katsura, Yoshimoto; Tasaka, Hiromichi; Komuro, Katsutoshi; Yamada, Takeshi; Asan, Toshihiko; Yamazaki, Akihiro; Yamazaki, Shudo
- SO Proceedings of the National Academy of Sciences of the United States of America, (1995) Vol. 92, No. 23, pp. 10693-10697.  
CODEN: PNASA6. ISSN: 0027-8424.
- DT Article
- LA English
- ED Entered STN: 12 Jan 1996  
Last Updated on STN: 12 Jan 1996
- AB A recombinant *Mycobacterium bovis* bacillus Calmette-Guerin (BCG) vector-based vaccine that secretes the V3 principal neutralizing epitope of human immunodeficiency virus (HIV) could induce immune response to the epitope and prevent the viral infection. By using the Japanese consensus sequence of HIV-1, we successfully constructed chimeric protein secretion vectors by selecting an appropriate insertion site of a carrier protein and established the principal neutralizing determinant (PND)-peptide secretion system in BCG. The recombinant BCG (rBCG)-inoculated guinea pigs were initially screened by delayed-type hypersensitivity (DTH) skin reactions to the PND peptide, followed by passive transfer of the DTH by the systemic route. Further, immunization of mice with the rBCG resulted in induction of cytotoxic T lymphocytes. The guinea pig immune antisera showed elevated titers to the PND peptide and neutralized HIV-MN, and administration of serum IgG from the vaccinated guinea pigs was effective in completely blocking the HIV infection in thymus/liver transplanted severe combined immunodeficiency (SCID)/hu or SCID/PBL mice. In addition, the immune serum IgG was shown to neutralize primary field isolates of HIV that match the neutralizing sequence motif by a peripheral blood mononuclear cell-based virus neutralization assay. The data support the idea that the antigen-secreting rBCG system can be used as a tool for development of HIV vaccines.
- L4 ANSWER 29 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
STN DUPLICATE 12
- AN 1995:157783 BIOSIS
- DN PREV199598172083
- TI Delayed-type hypersensitivity to a recombinant mycobacterial antigen, MPB64, in guinea pigs sensitized to *Mycobacterium tuberculosis* or *Mycobacterium bovis* BCG.
- AU Haga, Shinji; Yamaguchi, Ryuji; Nagai, Sadamu; Matsuo, Kazuhiro; Yamazaki, Akihiro; Nakamura, Reiko M. [Reprint author]
- CS 1-23-1, Toyama, Shinjuku-ku, Tokyo 162, Japan
- SO Journal of Leukocyte Biology, (1995) Vol. 57, No. 2, pp. 221-225.  
CODEN: JLBIE7. ISSN: 0741-5400.
- DT Article
- LA English
- ED Entered STN: 11 Apr 1995  
Last Updated on STN: 11 Apr 1995
- AB Recombinant MPB64 (rMPB64), a mycobacterial antigen, was obtained from an

Escherichia coli clone transformed with a recombinant expression vector, pMAL64c. The rMPB64 was examined for the activity to elicit delayed-type hypersensitivity (DTH) in guinea pigs injected with live Mycobacterium tuberculosis H37Rv or live M. bovis BCG Tokyo. It was found that rMPB64 has the same reactivity as native MPB64 (nMPB64) or MPT64 (nMPT64) and the potency to elicit DTH was 13.4 times higher than that of PPD. Because MPB64 is secreted only by living M. tuberculosis and some strains of BCG, it is possible to use this antigen for the diagnosis of tuberculosis.

L4 ANSWER 30 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
STN DUPLICATE 13  
AN 1994:110138 BIOSIS  
DN PREV199497123138  
TI Cytotoxic T lymphocyte response in mice induced by a recombinant  
BCG vaccination which produces an extracellular alpha antigen that  
fused with the human immunodeficiency virus type 1 envelope immunodominant  
domain in the V3 loop.  
AU Kameoka, Masanori; Nishino, Yoshii; Matsuo, Kazuhiro; Ohara,  
Naoya; Kimura, Takuro; Yamazaki, Akihiro; Yamada, Takeshi; Ikuta,  
Kazuyoshi [Reprint author]  
CS Section Serology, Institute Immunological Science, Hokkaido University,  
Kita-ku, Sapporo 060, Japan  
SO Vaccine, (1994) Vol. 12, No. 2, pp. 153-158.  
CODEN: VACCDE. ISSN: 0264-410X.  
DT Article  
LA English  
ED Entered STN: 14 Mar 1994  
Last Updated on STN: 14 Mar 1994  
AB The host immune response of cell-mediated immunity, particularly that of  
cytotoxic T lymphocytes (CTLs), is a major immune defence mechanism which  
may provide resistance to a human immunodeficiency virus type 1 (HIV-1)  
spread leading to acquired immune deficiency syndrome (AIDS). To prevent  
the accompanying activity of HIV-1 proteins responsible for the loss of  
helper T-lymphocyte function, it is crucial to develop a live attenuated  
recombinant vaccine expressing only T- or both T- and B-cell epitopes.  
Here, we examined the expression of the HIV-1 Env protein V3 region (15  
amino acids from Arg-315 to Lys-329) in Mycobacterium bovis BCG  
as a fused form with an extracellular alpha antigen of Mycobacterium  
kansasii. Balb/c mice inoculated with this recombinant BCG  
(rBCG), rapidly induced V3 peptide-specific CTLs. Target cell lysis was  
restricted to the murine class I major histocompatibility complex, H-2-d.  
A similar CTL response was also elicited after Balb/c mice were immunized  
with the same rBCG even when pre-inoculated with non-recombinant  
BCG. Thus, the rapid induction of HIV-1-specific CTLs indicates  
that this vaccine may be a therapeutic approach to preventing progression  
to AIDS.

L4 ANSWER 31 OF 47 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on  
STN DUPLICATE 14  
AN 1993:252168 BIOSIS  
DN PREV199395131343  
TI Cloning and sequencing of the gene for alpha antigen from Mycobacterium  
avium and mapping of B-cell epitopes.  
AU Ohara, Naoya [Reprint author]; Matsuo, Kazuhiro; Yamaguchi,  
Ryuji; Yamazaki, Akihiro; Tasaka, Hiromichi; Yamada, Takeshi  
CS Sch. Dentistry, Nagasaki University, Sakamoto 1-7-1, Nagasaki City 852,  
Japan  
SO Infection and Immunity, (1993) Vol. 61, No. 4, pp. 1173-1179.  
CODEN: INFIBR. ISSN: 0019-9567.  
DT Article  
LA English  
OS EMBL-X63437  
ED Entered STN: 21 May 1993

Last Updated on STN: 21 May 1993

AB The complete nucleotide sequence of alpha antigen secreted from *Mycobacterium avium* (A-alpha) was determined. The gene encodes 330 amino acids, including 40 amino acids for the signal peptide, followed by 290 amino acids for the mature protein with a molecular mass of 30,811 Da. This is the first sequence of A-alpha. Comparisons between A-alpha antigens of *Mycobacterium leprae*, *Mycobacterium bovis* BCG, and *Mycobacterium kansasii* showed highly homologous regions which suggested a conserved functional domain and two less homologous regions. Serological analysis of recombinant A-alpha, expressed by a series of deletion constructs, indicated the possibility that A-alpha carries at least six B-cell epitopes. The three antigenic determinants were common to *Mycobacterium tuberculosis*, *M. kansasii*, and *M. avium*. The results also suggested the possibility that there are three species-specific epitopes.

L4 ANSWER 32 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1991:201950 CAPLUS

DN 114:201950

TI Fusion protein based epitope mapping of the MPB57 protein from *Mycobacterium bovis* BCG and its epitope insertion into the native protein

AU Yamaguchi, Ryuji; Matsuo, Kazuhiro; Yamazaki, Akihiro; Kagawa, Hiroaki; Nagai, Sadamu; Yamada, Takeshi

CS Cent. Res. Lab., Ajinomoto Co. Inc., Kawasaki, 210, Japan

SO Canadian Journal of Microbiology (1991), 37(1), 7-13

CODEN: CJMIAZ; ISSN: 0008-4166

DT Journal

LA English

AB The gene coding for the 12-kDa protein (MPB57) of *M. bovis* BCG was recently cloned and sequenced. To map linear B-cell epitopes by  $\beta$ -galactosidase fusion proteins, convenient vectors (pUR278S, pUR288S, and pUR289S) were constructed with the SmaI site. Based on recognition by polyclonal antibodies, 2 epitope regions on the MPB57 protein were identified, both of which corresponded to the amino acid sequences Glu20 to Val45 (26 residues, epitope I region) and Ile78 to Leu86 (9 residues, epitope II). Complementary oligonucleotides encoding epitope II were synthesized, polymerized by a ligase reaction, inserted into the native MPB67 protein gene, and expressed in *Escherichia coli*, giving rise to epitope-inserted proteins. Their stability and potential uses are described.

L4 ANSWER 33 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 15

AN 1991:158537 CAPLUS

DN 114:158537

TI Cloning and expression of gene for antigen MPB70 of *Mycobacterium bovis* (BCG)

IN Yamada, Takeshi; Yamaguchi, Takashi; Matsuo, Kazuhiro; Yamazaki, Akihiro

PA Ajinomoto Co., Inc., Japan

SO Jpn. Kokai Tokkyo Koho, 11 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	JP 02195895	A	19900802	JP 1989-13270	19890124
PRAI	JP 1989-13270		19890124		

AB The gene encoding the BCG antigen MPB70 is cloned, sequenced, and its amino acid sequence deduced. Preparation of the MPB70 antigen as a fusion protein with the N-terminal region of human interleukin-2 is also shown. The antigen is useful in diagnosis of cattle-specific tuberculosis. The gene for antigen MPB70 was cloned from the chromosomal DNA library of the BCG (Tokyo strain) prepared in pUC19. The

signal sequence, the promoter region, and the Shine/Dalgarno sequence of the gene were also identified. Plasmid pT13SNco encoding the fusion protein from the trp promoter was provided and the chimeric gene expressed in *Escherichia coli*.

L4 ANSWER 34 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 16

AN 1991:581526 CAPLUS

DN 115:181526

TI Determinant of antigen MPB57 of *Mycobacterium bovis* BCG and its chemical or recombinant preparation

IN Yamaguchi, Takashi; Matsuo, Kazuhiro; Yamazaki, Akihiro; Yamada, Takeshi

PA Ajinomoto Co., Inc., Japan

SO Jpn. Kokai Tokkyo Koho, 9 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	JP 02169599	A	19900629	JP 1988-322515	19881221
PRAI	JP 1988-322515		19881221		

AB A synthetic gene encoding a determinant of the MPB57 antigen of *Mycobacterium bovis* BCG (BCG) is cloned and expressed as a polymer in *Escherichia coli*. The recombinant determinant can be used for diagnosis of tuberculosis. The Sau 3AI fragment of plasmid pKKM 57 carrying a synthetic gene encoding the MPB57 antigen of BCG was used to prepare an expression library in plasmids pUR 278, 288, and 289. Polyclonal antibody to MPB57 was used to screen for the gene encoding the determinant and its amino acid sequence deduced. Plasmid pKKM 57(A) carrying a synthetic gene encoding the polymeric antigen determinant of the MPB 57 protein and prepared and expressed in *E. coli*.

L4 ANSWER 35 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 17

AN 1990:435890 CAPLUS

DN 113:35890

TI Cloning and expression of antigen MPB57 protein gene of *Mycobacterium bovis* BCG

IN Yamaguchi, Takashi; Matsuo, Kazuhiro; Yamazaki, Akihiro; Yamada, Takeshi

PA Ajinomoto Co., Inc., Japan

SO Jpn. Kokai Tokkyo Koho, 10 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	JP 02053496	A	19900222	JP 1988-205444	19880818
PRAI	JP 1988-205444		19880818		

AB The antigen MPB57 protein gene of *M. bovis* BCG was cloned and expressed in *Escherichia coli*. The antigen gene was cloned on the vector plasmid pKK233-2 to generate the recombinant plasmid pKKM57. The sequence of the gene is given.

L4 ANSWER 36 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1991:242070 CAPLUS

DN 114:242070

TI *Mycobacterium kansasii*  $\alpha$  antigen gene cloning and expression

IN Matsuo, Kazuhiro; Yamaguchi, Takashi; Yamazaki, Akihiro; Yamada, Takeshi

PA Ajinomoto Co., Inc., Japan

SO Jpn. Kokai Tokkyo Koho, 11 pp.

CODEN: JKXXAF

DT Patent  
LA Japanese  
FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	JP 02308793	A	19901221	JP 1989-128091	19890522
PRAI	JP 1989-128091		19890522		

AB  $\alpha$ -Antigen of of M. kansasii is cloned and expressed in Escherichia coli. The antigen is useful for the diagnosis of tuberculosis-like, M. kansasii-associated respiratory diseases. The gene was cloned from a genomic library of M. kansasii ATCC12478 on pUC18 using a fragment of  $\alpha$ -antigen gene of BCG bacteria as probe. Plasmid pUCK201 encoding the  $\alpha$ -antigen was constructed and transformed into E. coli. By aerobic fermentation, the E. coli transformants produced and secreted  $\alpha$ -antigen (mol. weight, 30000).

L4 ANSWER 37 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN  
AN 1991:466177 CAPLUS  
DN 115:66177

TI Mycobacterial secretory expression vectors and transformants for use as live vaccine

IN Matsuo, Kazuhiro; Yamaguchi, Ryuji; Yamazaki, Akihiro; Yamada, Takeshi

PA Ajinomoto Co., Inc., Japan

SO Eur. Pat. Appl., 18 pp.

CODEN: EPXXDW

DT Patent  
LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	EP 400973	A1	19901205	EP 1990-305849	19900530
	EP 400973	B1	19960731		
	R: DE, FR, GB				
	JP 03072888	A	19910328	JP 1990-64310	19900316
	JP 2903414	B2	19990607		
	US 6015696	A	20000118	US 1994-193899	19940209
PRAI	JP 1989-135855	A	19890531		
	JP 1990-64310	A	19900316		
	US 1990-531448	B1	19900531		

AB Secretory expression vectors for the expression of heterologous genes in mycobacteria, especially Mycobacterium bovis BCG (BCG), are prepared. Mycobacteria harboring the expression vectors and producing and secreting the heterologous proteins are useful as live vaccines. The use of recombinant mycobacteria as live vaccines is safe and also has the following advantages: long-lasting activity, strong antigenicity because of the strong adjuvant activity of BCG, and low cost. The expression of gene for  $\alpha$ -antigen of M. kansasii,  $\beta$ -lactamase of Escherichia coli, and a fusion protein of  $\alpha$ -antigen and the B cell epitope of HIV-1 gag protein p17 protein were shown. All the described proteins were successfully secreted into the culture medium.

L4 ANSWER 38 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN  
AN 1991:22059 CAPLUS  
DN 114:22059

TI Establishment of a foreign antigen secretion system in mycobacteria

AU Matsuo, Kazuhiro; Yamaguchi, Ryuji; Yamazaki, Akihiro; Tasaka, Hiromichi; Terasaka, Kunihiro; Totsuka, Masayoshi; Kobayashi, Koumei; Yukitake, Hideharu; Yamada, Takeshi

CS Cent. Res. Lab., Ajinomoto Co. Inc., Kawasaki, 210, Japan

SO Infection and Immunity (1990), 58(12), 4049-54

CODEN: INFIBR; ISSN: 0019-9567

DT Journal  
LA English

AB In order to develop recombinant Mycobacterium bovis BCG into a useful multivaccine vehicle, a foreign antigen secretion system was established in mycobacteria in which an extracellular  $\alpha$  antigen of M. kansasii was utilized as a carrier. By using this system, a B-cell epitope (Glu-12-Leu-Asp-Arg-Trp-Glu-Lys-Ile-19) of human immunodeficiency virus type 1 p17gag, which was identified by a fusion protein-based method, has been successfully obtained from BCG along with the  $\alpha$  antigen. This is the first report of expression and secretion of a foreign viral antigen from BCG. It is possible that the system can become a universal vaccination vehicle applicable to protection against various infectious diseases.

L4 ANSWER 39 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1990:453246 CAPLUS

DN 113:53246

TI Cloning and expression of the gene for the cross-reactive  $\alpha$  antigen of Mycobacterium kansasii

AU Matsuo, Kazuhiro; Yamaguchi, Ryuji; Yamazaki, Akihiro; Tasaka, Hiromichi; Terasaka, Kunihiro; Yamada, Takeshi

CS Cent. Res. Lab., Ajinomoto Co. Inc., Kawasaki, 210, Japan

SO Infection and Immunity (1990), 58(2), 550-6

CODEN: INFIBR; ISSN: 0019-9567

DT Journal

LA English

AB The gene for the extracellular  $\alpha$  antigen of M. kansasii was cloned by using the  $\alpha$ -antigen gene fragments of M. bovis BCG as probes. Gene anal. revealed that this gene encodes 325 amino acid residues, including 40 aa for the signal peptide, followed by 285 aa for the mature protein. A comparison of the nucleotide sequences of the genes isolated from these 2 mycobacterial species showed that the levels of DNA and aa homol. were 84.8 and 89.1%, resp. The hydropathy profiles were also compared, and 2 highly changed hydrophilic regions were observed, which might account for the antigenic diversity of this antigen or its acquirement of antigenic specificity.

L4 ANSWER 40 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 18

AN 1990:453702 CAPLUS

DN 113:53702

TI Cloning of gene for MPB 64 protein of Mycobacterium bovis BCG

IN Yamaguchi, Takashi; Matsuo, Kazuhiro; Yamazaki, Akihiro; Abe, Choji; Nagai, Sadamu; Yamada, Takeshi

PA Ajinomoto Co., Inc., Japan

SO Jpn. Kokai Tokkyo Koho, 12 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	JP 01247094	A	19891002	JP 1988-77366	19880330
PRAI	JP 1988-77366		19880330		

AB The gene encoding protein MPB64 of M. bovis BCG, a protein that is useful for diagnosis of tuberculosis, is cloned and its amino acid sequence deduced. An Escherichia coli expression plasmid, pKKM64, is also provided and used for expression of the gene.

L4 ANSWER 41 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 19

AN 1990:153072 CAPLUS

DN 112:153072

TI Molecular cloning and expression of the gene for the  $\alpha$ -antigen of Mycobacterium bovis BCG

IN Matsuo, Kazuhiro; Yamaguchi, Takashi; Yamazaki, Akihiro; Yamada, Takeshi

PA Ajinomoto Co., Inc., Japan

SO Jpn. Kokai Tokkyo Koho, 26 pp.

CODEN: JKXXAF

DT Patent

LA Japanese

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	JP 01144994	A	19890607	JP 1987-305250	19871202
PRAI	JP 1987-305250		19871202		

AB The gene encoding  $\alpha$ -antigen of Mycobacterium bovis BCG, that can be used in manufacturing tuberculin, is cloned and sequenced. An Escherichia coli expression plasmid is also provided. Plasmid p $\alpha$ L-1 containing the gene encoding the  $\alpha$ -antigen of M. bovis BCG Tokyo was found by screening of a chromosomal DNA library and the DNA sequence of the gene determined. Expression plasmid pKK $\alpha$ -2 was then constructed and its gene product was identified.

L4 ANSWER 42 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1989:568195 CAPLUS

DN 111:168195

TI Cloning and characterization of the gene for immunogenic protein MPB64 of Mycobacterium bovis BCG

AU Yamaguchi, Ryuji; Matsuo, Kazuhiro; Yamazaki, Akihiro; Abe, Chiyoji; Nagai, Sadamu; Terasaka, Kunihiro; Yamada, Takeshi

CS Cent. Res. Lab., Ajinomoto Co., Inc., Kawasaki, 210, Japan

SO Infection and Immunity (1989), 57(1), 283-8

CODEN: INFIBR; ISSN: 0019-9567

DT Journal

LA English

AB The gene for immunogenic protein MPB64 found in culture filtrates of only Mycobacterium tuberculosis and some strains of M. bovis BCG was cloned by using a single-probe method and was sequenced. The gene anal. revealed that the structural gene for MPB64 consisted of 618 base pairs, and its deduced mol. weight was 22,400. Twenty-two amino acids for a putative signal peptide and 205 amino acids for the MPB64 protein were observed. In the coding region, the third letter of the codon showed a biased codon and a high G+C content (78.5%). The gene was expressed in Escherichia coli by using an E. coli expression vector. The product showed migration similar to that of the authentic MPB64 protein by electrophoresis and reacted with the polyclonal and the monoclonal antibodies raised against the MPB64 protein. The strict specificity of MPB64 could be applied to immunodiagnosis of tuberculosis.

L4 ANSWER 43 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1989:592690 CAPLUS

DN 111:192690

TI Complete nucleotide sequence of immunogenic protein MPB70 from Mycobacterium bovis BCG

AU Terasaka, Kunihiro; Yamaguchi, Ryuji; Matsuo, Kazuhiro; Yamazaki, Akihiro; Nagai, Sadamu; Yamada, Takeshi

CS Res. Inst. Microb. Dis., Osaka Univ., Osaka, 565, Japan

SO FEMS Microbiology Letters (1989), 58(2-3), 273-6

CODEN: FMLED7; ISSN: 0378-1097

DT Journal

LA English

AB The extracellular protein MPB70 is a heat-stable immunogenic protein which was found in the culture filtrate of M. bovis BCG Japanese. The authors determined the complete nucleotide and amino acid sequences of MPB70. The N-terminal sequence revealed that the signal peptide (SP) consisted of 30 amino acids and that the mature protein had 163 amino acids with a mol. weight of 16,305. The SP displayed a characteristic feature of an Ala-rich property which would be efficient in a SP function.

L4 ANSWER 44 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN

AN 1989:148838 CAPLUS  
 DN 110:148838  
 TI Cloning and expression of the Mycobacterium bovis BCG gene for extracellular  $\alpha$  antigen  
 AU Matsuo, Kazuhiro; Yamaguchi, Ryuji; Yamazaki, Akihiro; Tasaka, Hiromichi; Yamada, Takeshi  
 CS Cent. Res. Lab., Ajinomoto Co. Inc., Kawasaki, 210, Japan  
 SO Journal of Bacteriology (1988), 170(9), 3847-54  
 CODEN: JOBAAY; ISSN: 0021-9193  
 DT Journal  
 LA English  
 AB The gene for the extracellular  $\alpha$  antigen of M. bovis BCG was cloned by using a single probe restricted to G or C in the third position. This technique should have great potential for the isolation of mycobacterial antigen genes. The gene anal. revealed that the  $\alpha$  antigen gene encoded 323 amino acid residues, including 40 amino acids for signal peptide followed by 283 amino acids for mature protein. This is the first report on the structure of the mycobacterial signal peptide. The promoter-like sequence and ribosome-binding site were observed upstream of the open reading frame. In the coding region, the third position of the codon showed high G + C content (86%). The gene was expressed as an unfused protein in Escherichia coli by using an E. coli expression vector. This protein, which reacted with polyclonal antibody raised  $\alpha$  antigen from M. tuberculosis, would be applicable to the immunodiagnosis of tuberculosis.

L4 ANSWER 45 OF 47 CAPLUS COPYRIGHT 2007 ACS on STN  
 AN 1989:491323 CAPLUS  
 DN 111:91323  
 TI Immunogenic protein MPB57 from Mycobacterium bovis BCG: molecular cloning, nucleotide sequence and expression  
 AU Yamaguchi, Ryuji; Matsuo, Kazuhiro; Yamazaki, Akihiro; Nagai, Sadamu; Terasaka, Kunihiro; Yamada, Takeshi  
 CS Cent. Res. Lab., Ajinomoto Co. Inc., Kawasaki, 210, Japan  
 SO FEBS Letters (1988), 240(1-2), 115-17  
 CODEN: FEBLAL; ISSN: 0014-5793  
 DT Journal  
 LA English  
 AB A single-probe method was used to clone the gene for an immunogenic MPB57 protein of M. bovis BCG. The nucleotide sequence includes an ORF of 300 base pairs encoding a protein of 99 amino acids with an Mr of 10,818. This cloned gene was expressed in an Escherichia coli expression vector to give a mature protein which reacted with a polyclonal antibody raised against MPB57.

L4 ANSWER 46 OF 47 JAPIO (C) 2007 JPO on STN  
 AN 2006-149234 JAPIO  
 TI PRIME-BOOST VACCINATION METHOD  
 IN HONDA MITSUO; MATSUO KAZUHIRO; HAMANO RYUICHI; IZUMI YASUYUKI; PROMKHATKAEW DUANTHANORM; BALACHANDRA KRUAUVON; SUTTHENT RUENGPUNG  
 PA JAPAN SCIENCE & TECHNOLOGY AGENCY  
 THAILAND MINISTRY OF PUBLIC HEALTH DEPARTMENT OF MEDICAL SCIENCES  
 NATIONAL INSTITUTE OF INFECTIOUS DISEASES  
 PI JP 2006149234 A 20060615 Heisei  
 AI JP 2004-341283 (JP2004341283 Heisei) 20041125  
 PRAI JP 2004-341283 20041125  
 SO PATENT ABSTRACTS OF JAPAN (CD-ROM), Unexamined Applications, Vol. 2006  
 AB PROBLEM TO BE SOLVED: To provide a new vaccination strategy to HIV-1 CEF01AE.  
 SOLUTION: The prime-boost vaccination method comprises a priming step with a recombinant BCG vaccine and one or more boosting steps with the recombinant vaccine. Both of the recombinant BCG vaccine for priming step and the recombinant vaccine for boosting step contain at least one gene of HIV-1 CRF01AE strain.



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L4 ANSWER 47 OF 47 JAPIO (C) 2007 JPO on STN  
AN 2002-068990 JAPIO  
TI Th1-TYPE CYTOKINE ACTIVATOR, METHOD FOR PRODUCING THE SAME, ATOPIC DISEASE  
PROPHYLACTIC/THERAPEUTIC VACCINE AND METHOD FOR PRODUCING THE VACCINE, AND  
METHOD FOR CONVERSION INTO Th1-TYPE CYTOKINE-DOMINANT CONDITION  
IN YASUTOMI YASUHIRO; MIZUTANI HITOSHI; MATSUO KAZUHIRO  
PA JAPAN BCG SEIZO KK  
YASUTOMI YASUHIRO  
MIZUTANI HITOSHI  
MATSUO KAZUHIRO  
PI JP 2002068990 A 20020308 Heisei  
AI JP 2000-261016 (JP2000261016 Heisei) 20000830  
PRAI JP 2000-261016 20000830  
SO PATENT ABSTRACTS OF JAPAN (CD-ROM), Unexamined Applications, Vol. 2002  
AB PROBLEM TO BE SOLVED: To provide a vaccine functioning to bring a  
Th2-dominant cytokine condition to Th1-dominant cytokine condition, to  
provide a method for producing the vaccine, and to provide a therapeutic  
method using the vaccine so as to contribute to treating and preventing  
various atopic diseases.  
SOLUTION: The objective Th1-type cytokine activator is characterized by  
mainly comprising one kind among BCG heat-killed bacterium and  
its microbial cell destruction products. The method for producing the  
Th1-type cytokine activator is provided. The therapeutic method using the  
vaccine as the activator is provided.  
COPYRIGHT: (C)2002,JPO

=> e kanekiyo masaru/au

E1 12 KANEKIYO MASAHIRO/AU  
E2 16 KANEKIYO MASAHO/AU  
E3 10 --> KANEKIYO MASARU/AU  
E4 1 KANEKIYO MASATO/AU  
E5 10 KANEKIYO MITSUGI/AU  
E6 4 KANEKIYO MITSUGU/AU  
E7 3 KANEKIYO N/AU  
E8 2 KANEKIYO NAKO/AU  
E9 9 KANEKIYO NOBORU/AU  
E10 3 KANEKIYO NORIHI/AU  
E11 2 KANEKIYO NORIFUKU/AU  
E12 2 KANEKIYO NORIHI/AU

=> s e3

L5 10 "KANEKIYO MASARU"/AU

=> dup rem l5

PROCESSING COMPLETED FOR L5

L6 5 DUP REM L5 (5 DUPLICATES REMOVED)

=> d bib ab 1-

YOU HAVE REQUESTED DATA FROM 5 ANSWERS - CONTINUE? Y/(N):y

L6 ANSWER 1 OF 5 USPATFULL on STN  
AN 2006:247197 USPATFULL  
TI Recombination bcg vaccine  
IN Honda, Mitsuo, Tokyo, JAPAN  
Matsuo, Kazuhiro, Kanagawa, JAPAN  
Kanekiyo, Masaru, Tokyo, JAPAN  
PI US 2006210586 A1 20060921  
AI US 2003-524586 A1 20030813 (10)  
WO 2003-JP10303 20030813  
20050331 PCT 371 date  
PRAI JP 2002-237610 20020816

DT Utility  
 FS APPLICATION  
 LREP WENDEROTH, LIND & PONACK, L.L.P., 2033 K STREET N. W., SUITE 800,  
 WASHINGTON, DC, 20006-1021, US  
 CLMN Number of Claims: 2  
 ECL Exemplary Claim: 1  
 DRWN 5 Drawing Page(s)  
 LN.CNT 640  
 CAS INDEXING IS AVAILABLE FOR THIS PATENT.  
 AB A recombinant BCG vaccine being transformed with an expression vector  
 that has a polynucleotide encoding a foreign antigenic protein, wherein  
 the polynucleotide is a modified one in which a third position of each  
 codon is substituted with G or C without a change of an amino acid. This  
 recombinant BCG vaccine has an excellent expression rate of antigenic  
 protein and, as a result, capable of inducing a sufficient immune  
 response against target infectious disease, cancer, or the like at the  
 same dose as that of the typical BCG vaccine.

L6 ANSWER 2 OF 5 CAPLUS COPYRIGHT 2007 ACS on STN  
 AN 2006:1086167 CAPLUS  
 DN 145:453348  
 TI Priming-boosting vaccination with recombinant Mycobacterium bovis bacillus  
 Calmette-Guerin and a nonreplicating vaccinia virus recombinant leads to  
 long-lasting and effective immunity. [Erratum to document cited in  
 CA143:365230]  
 AU Ami, Yasushi; Izumi, Yasuyuki; Matsuo, Kazuhiro; Someya, Kenji;  
 Kanekiyo, Masaru; Horibata, Shigeo; Yoshino, Naoto; Sakai, Koji;  
 Shinohara, Katsuaki; Matsumoto, Sohkiichi; Yamada, Takeshi; Yamazaki,  
 Shudo; Yamamoto, Naoki; Honda, Mitsuo  
 CS Division of Experimental Animal Research, AIDS Research Center, Division  
 of Biosafety Control and Research, National Institute of Infectious  
 Diseases, Toyama 1-23-1, Shinjuku-ku, Tokyo, 162-8640, Japan  
 SO Journal of Virology (2006), 80(20), 10288  
 CODEN: JOVIAM; ISSN: 0022-538X  
 PB American Society for Microbiology  
 DT Journal  
 LA English  
 AB On page 12873, Table 1, in groups 1, 2 and 3, rBCG was primed at week 3,  
 followed by booster immunization of rDIs at weeks 50 and 54. In the same  
 table, groups 4 and 5, rDIs was primed at weeks 0 and 12, followed by  
 booster immunization of rBCG at week 50. Then all animals were challenged  
 with virulent SHIV KS661c at week 57. On page 12874, Figure 2: The weeks  
 after immunization shown on the x axis in panel A should read: "3, 7, 11,  
 19, 27, 35, 50, 53, 54, and 56.". The weeks after immunization shown on  
 the x axis in panel B should read: "0, 4, 8, 16, 24, 32, 50, 52, 53, 54,  
 and 56.". Although the patterns and magnitudes of the kinetics were  
 almost the same as the original ones, the standard deviation of the ELISPOT  
 data at the peak response at 54 wk after immunization was 500, which was  
 five times more than originally reported. Spot-forming cells were counted  
 by using a KS ELISPOT system after 35 wk of immunization. Before that, we  
 counted SFCs using an inverted microscope. On page 12878,  
 Acknowledgments, paragraph 1, the last sentence of the paragraph should be  
 deleted. These changes do not alter the conclusions of the article.

L6 ANSWER 3 OF 5 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN  
 DUPLICATE 1  
 AN 2006:8301 BIOSIS  
 DN PREV200600008996  
 TI Priming-boosting vaccination with recombinant Mycobacterium bovis bacillus  
 Calmette-Guerin and a nonreplicating vaccinia virus recombinant leads to  
 long-lasting and effective immunity.  
 AU Ami, Yasushi; Izumi, Yasuyuki; Matsuo, Kazuhiro; Someya, Kenji;  
 Kanekiyo, Masaru; Horibata, Shigeo; Yoshino, Naoto; Sakai, Koji;  
 Shinohara, Katsuaki; Matsumoto, Sohkiichi; Yamada, Takeshi; Yamazaki,

Shudo; Yamamoto, Naoki; Honda, Mitsuo [Reprint Author]  
 CS Natl Inst Infect Dis, Ctr AIDS Res, Shinjuku Ku, Toyama 1-23-1, Tokyo  
 1628640, Japan  
 mhonda@nih.go.jp  
 SO Journal of Virology, (OCT 2005) Vol. 79, No. 20, pp. 12871-12879.  
 CODEN: JOVIAM. ISSN: 0022-538X.  
 DT Article  
 LA English  
 ED Entered STN: 14 Dec 2005  
 Last Updated on STN: 14 Dec 2005  
 AB Virus-specific T-cell responses can limit immunodeficiency virus type 1  
 (HIV-1) transmission and prevent disease progression and so could serve as  
 the basis for an affordable, safe, and effective vaccine in humans. To  
 assess their potential for a vaccine, we used Mycobacterium bovis bacillus  
 Calmette-Guerin (BCG)-Tokyo and a replication-deficient vaccinia virus  
 strain (DIs) as vectors to express full-length gag from simian  
 immunodeficiency viruses (SIVs) (rBCG-SIVgag and rDIsSIVgag). Cynomolgus  
 macaques were vaccinated with either rBCG-SIVgag dermally as a single  
 modality or in combination with rDIsSIVgag intravenously. When  
 cynomolgus macaques were primed with rBCG-SIVgag and then boosted with  
 rDIsSIVgag, high levels of gamma interferon (IFN-gamma) spot-forming cells  
 specific for SIV Gag were induced. This combination regimen elicited  
 effective protective immunity against mucosal challenge with pathogenic  
 simian-human immunodeficiency virus for the 1 year the macaques were under  
 observation. Antigen-specific intracellular IFN-gamma activity was  
 similarly induced in each of the macaques with the priming-boosting  
 regimen. Other groups receiving the opposite combination or the  
 single-modality vaccines were not effectively protected. These results  
 suggest that a recombinant M. bovis BCG-based vector may have potential as  
 an HIV/AIDS vaccine when administered in combination with a  
 replication-deficient vaccinia virus DIs vector in a priming-boosting  
 strategy.

L6 ANSWER 4 OF 5 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN  
 DUPLICATE 2  
 AN 2005:399376 BIOSIS  
 DN PREV200510190449  
 TI Mycobacterial codon optimization enhances antigen expression and  
 virus-specific immune responses in recombinant Mycobacterium bovis bacille  
 Calmette-Guerin expressing human immunodeficiency virus type 1 Gag.  
 AU Kanekiyo, Masaru; Matsuo, Kazuhiro; Hamatake, Makiko; Hamano,  
 Takaichi; Ohsu, Takeaki; Matsumoto, Sohkiichi; Yamada, Takeshi; Yamazaki,  
 Shudo; Hasegawa, Atsuhiko; Yamamoto, Naoki; Honda, Mitsuo [Reprint Author]  
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 SO Journal of Virology, (JUL 2005) Vol. 79, No. 14, pp. 8716-8723.  
 CODEN: JOVIAM. ISSN: 0022-538X.  
 DT Article  
 LA English  
 ED Entered STN: 5 Oct 2005  
 Last Updated on STN: 5 Oct 2005  
 AB Although its potential for vaccine development is already known, the  
 introduction of recombinant human immunodeficiency virus (HIV) genes to  
 Mycobacterium bovis bacille Calmette-Guerin (BCG) has thus far elicited  
 only limited responses. In order to improve the expression levels, we  
 optimized the codon usage of the HIV type 1 (HIV-1) p24 antigen gene of  
 gag (p24 gag) and established a codon-optimized recombinant BCG (rBCG)-p24  
 Gag which expressed a 40-fold-higher level of p24 Gag than did that of  
 nonoptimized rBCG-p24 Gag. Inoculation of mice with the codon-optimized  
 rBCG-p24 Gag elicited effective immunity, as evidenced by virus-specific  
 lymphocyte proliferation, gamma interferon ELISPOT cell induction, and  
 antibody production. In contrast, inoculation of animals with the  
 nonoptimized rBCG-p24 Gag induced only low levels of immune responses.

Furthermore, a dose as small as 0.01 mg of the codon-optimized rBCG per animal proved capable of eliciting immune responses, suggesting that even low doses of a codon-optimized rBCG-based vaccine could effectively elicit HIV-1-specific immune responses.

L6 ANSWER 5 OF 5 CAPLUS COPYRIGHT 2007 ACS on STN  
 AN 2004:162604 CAPLUS  
 DN 140:216160  
 TI Recombinant BCG vaccines against infection, cancer and other diseases  
 IN Honda, Mitsuo; Matsuo, Kazuhiro; Kanekiyo, Masaru  
 PA Japan Science and Technology Corporation, Japan; Japan as Represented by  
 Director General of National Institute of Infectious Diseases  
 SO PCT Int. Appl., 31 pp.  
 CODEN: PIXXD2  
 DT Patent  
 LA Japanese  
 FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
PI	WO 2004016280	A1	20040226	WO 2003-JP10303	20030813
	W: IN, JP, US				
	RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR				
	EP 1535627	A1	20050601	EP 2003-788098	20030813
	R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, FI, RO, CY, TR, BG, CZ, EE, HU, SK				
	US 2006210586	A1	20060921	US 2005-524586	20050331
PRAI	JP 2002-237610	A	20020816		
	WO 2003-JP10303	W	20030813		
AB	A recombinant BCG vaccine obtained by transformation with an expression vector carrying a polynucleotide coding for an ecdemic antigenic protein, which recombinant BCG vaccine consists of a modified type polynucleotide comprising a polynucleotide having the third base of each of the codons thereof substituted with G or C without changing of the type of amino acid. This recombinant BCG vaccine excels in the amount of antigenic protein expressed, so that even with the same dosage as employed for conventional BCG vaccines, the recombinant BCG vaccine can induce satisfactory immune response to target infectious diseases, cancer, etc. In example, recombinant BCG comprising hsp60 gene and HIV-1 gag p24 gene was prepared for use as vaccine.				

RE.CNT 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD  
 ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> s (recombinant BCG)and(expression vector?)and (high g)

L7 3 (RECOMBINANT BCG) AND(EXPRESSION VECTOR?) AND (HIGH G)

=> dup rem 17

PROCESSING COMPLETED FOR L7

L8 3 DUP REM L7 (0 DUPLICATES REMOVED)

=> d bib ab 1-

YOU HAVE REQUESTED DATA FROM 3 ANSWERS - CONTINUE? Y/(N):y

L8 ANSWER 1 OF 3 USPATFULL on STN  
 AN 2006:247197 USPATFULL  
 TI Recombination bcg vaccine  
 IN Honda, Mitsuo, Tokyo, JAPAN  
 Matsuo, Kazuhiro, Kanagawa, JAPAN  
 Kanekiyo, Masaru, Tokyo, JAPAN  
 PI US 2006210586 A1 20060921  
 AI US 2003-524586 A1 20030813 (10)  
 WO 2003-JP10303 20030813  
 20050331 PCT 371 date

PRAI JP 2002-237610 20020816

DT Utility

FS APPLICATION

LREP WENDEROTH, LIND & PONACK, L.L.P., 2033 K STREET N. W., SUITE 800,  
WASHINGTON, DC, 20006-1021, US

CLMN Number of Claims: 2

ECL Exemplary Claim: 1

DRWN 5 Drawing Page(s)

LN.CNT 640

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB A recombinant BCG vaccine being transformed with an expression vector that has a polynucleotide encoding a foreign antigenic protein, wherein the polynucleotide is a modified one in which a third position of each codon is substituted with G or C without a change of an amino acid. This recombinant BCG vaccine has an excellent expression rate of antigenic protein and, as a result, capable of inducing a sufficient immune response against target infectious disease, cancer, or the like at the same dose as that of the typical BCG vaccine.

L8 ANSWER 2 OF 3 USPATFULL on STN

AN 2005:254317 USPATFULL

TI Identification of virulence associated regions rd1 and rd5 leading to improve vaccine of m. bovis bcg and m. microti

IN Cole, Stewart, Clamart, FRANCE

Pym, Alexander S., London, UNITED KINGDOM

Brosch, Roland, Paris, FRANCE

Brodin, Priscille, Paris, FRANCE

Majlessi, Laleh, Montigny Le Bretonneux, FRANCE

Demangel, Caroline, Paris, FRANCE

Leclerc, Claude, Paris, FRANCE

PI US 2005220811 A1 20051006

AI US 2003-510021 A1 20030401 (10)

WO 2003-IB1789 20030401

20041001 PCT 371 date

PRAI EP 2002-290864 20020405

DT Utility

FS APPLICATION

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AVENUE, NW, WASHINGTON, DC, 20001-4413, US

CLMN Number of Claims: 59

ECL Exemplary Claim: 1

DRWN 16 Drawing Page(s)

LN.CNT 5548

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention relates to a strain of M. bovis BCG or M. microti, wherein said strain has integrated part or all of the RD1 region responsible for enhanced immunogenicity of the tubercle bacilli, especially the ESAT-6 and CFP-10 genes. These strains will be referred as the M bovis BCG::RD1 or M. microti::RD1 strains and are useful as a new improved vaccine for preventing tuberculosis and as a therapeutical product enhancing the stimulation of the immune system for the treatment of bladder cancer. These strains are also useful for the expression and presentation of heterologous antigens and molecule that are of therapeutic or prophylactic interest.

L8 ANSWER 3 OF 3 USPATFULL on STN

AN 2005:243031 USPATFULL

TI Identification and cloning of a mycobacterial antigen corresponding to a heparin-binding haemagglutinin

IN Menozzi, Franco, Mons-Hyon, BELGIUM

Locht, Camille, Wannehain, FRANCE

PA Institut National de la Sante et de la Recherche Medicale, Paris, FRANCE  
(non-U.S. corporation)

Institut Pasteur de Lille, Lille, FRANCE (non-U.S. corporation)  
PI US 6949345 B1 20050927  
AI US 1998-192579 19981117 (9)  
RLI Continuation of Ser. No. WO 1997-FR886, filed on 20 May 1997, PENDING  
PRAI FR 1996-6168 19960517  
DT Utility  
FS GRANTED  
EXNAM Primary Examiner: Swartz, Rodney P  
LREP Nixon & Vanderhye P.C.  
CLMN Number of Claims: 3  
ECL Exemplary Claim: 1  
DRWN 13 Drawing Figure(s); 11 Drawing Page(s)  
LN.CNT 1104  
CAS INDEXING IS AVAILABLE FOR THIS PATENT.

AB The present invention relates to peptide sequences enabling mycobacteria to adhere to host cells (e.g., epithelial cells). More particularly, the invention relates to a mycobacterial heparin-binding haemagglutinin type antigen from *M. bovis* ECG or *M. tuberculosis*. The invention also relates to a recombinant peptide sequence enabling mycobacteria to adhere to host cells. The polypeptides can be used to prepare vaccines against mycobacterial infections and for serological diagnosis of mycobacterial infections.

=> d kwic 3

L8 ANSWER 3 OF 3 USPATFULL on STN

SUMM . . . its genome. In one preferred embodiment of the invention, the recombinant host cell is BCG, but not exclusively, for which expression vectors directly usable for developing recombinant BCG for use in man or animal have been developed.

DETD The sequence of two pairs of oligonucleotides was derived from the internal HBHA peptide sequences. The generally high G +C content in the mycobacterial DNA has led the inventors to favour G or C in the third position of the. . .

DETD . . . was purified by electro-elution after migration in a 1% agarose gel using standard procedures (17) and finally cloned in the expression vector pKK388-1 (Clontech, Palo Alto, Calif., UA) previously restricted with NcoI and KpnI. The recombinant plasmid was then introduced into E. . .

DETD . . . Coomassie blue shows the synthesis of a polypeptide of about 27 kDa in *E. coli* XL-1 Blue carrying the HBHA expression vector and derepressed by IPTG. This polypeptide, recognised in immunoblotting by a murine antiserum directed against the purified HBHA protein of. . . carrying the pKK388-1 vector with no insert, showing that its synthesis depends on the cloned BCG DNA sequence in the expression vector. As this is also the sequence coding for the BCG HBHA, it was surprising to observe that the recombinant polypeptide. . .